

Conference on  
**BIOLOGICAL INFORMATICS**

6-8 July 1998

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Australian Academy of Science, Canberra, Australia

# **What is Bioinformatics?**

( <http://www.esp.org/rjr/canberra.pdf> )

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## Abstract

In the last 25 years, Moore's Law has transformed society, delivering exponentially better computers at exponentially lower prices. Bioinformatics is the application of powerful, affordable information technology to the problems of biology. With \$2500 desktop PCs now delivering more raw computing power than the first Cray, bioinformatics is rapidly becoming the critical technology for 21st Century biology.

DNA is legitimately seen as a biological mass-storage device, making bioinformatics a *sine qua non* for genomic research. Others areas of biological investigation are equally information rich — an exhaustive tabulation of the Earth's biodiversity would involve a cross-index of the millions of known species against the approximately 500,000,000,000,000 square meters of the Earth's surface.

Bioinformatics is also becoming a scholarly discipline in its own right, melding information science with computer science, seasoning it with engineering methods, and applying it to the most information rich component of the known universe — the Biosphere.

# What is Bioinformatics?

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Bioinformatics is:

- the use of computers in pursuit of biological research.
- an emerging new discipline, with its own goals, research program, and practitioners.
- the *sine qua non* for 21st Century biology.
- all of the above.

# Topics

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- Bioinformatics is emerging as an independent discipline.

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- Bioinformatics is emerging as an independent discipline.
- **A connected, federated information infrastructure for biology is needed.**

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- Bioinformatics is emerging as an independent discipline.
- A connected, federated information infrastructure for biology is needed.
- **Current support for public bio-information infrastructure seems inadequate.**

# Introduction

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## *Magical Technology*

# Magic

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To a person from 1897, much current technology would seem like magic.

# Magic

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# Magic

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What technology of 2097 would seem magical to a person from 1997?

**Candidate:** Biotechnology so advanced that the distinction between living and non-living is blurred.

Information technology so advanced that access to information is immediate and universal.



# Moore's Law

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*Transforms InfoTech  
(and everything else)*

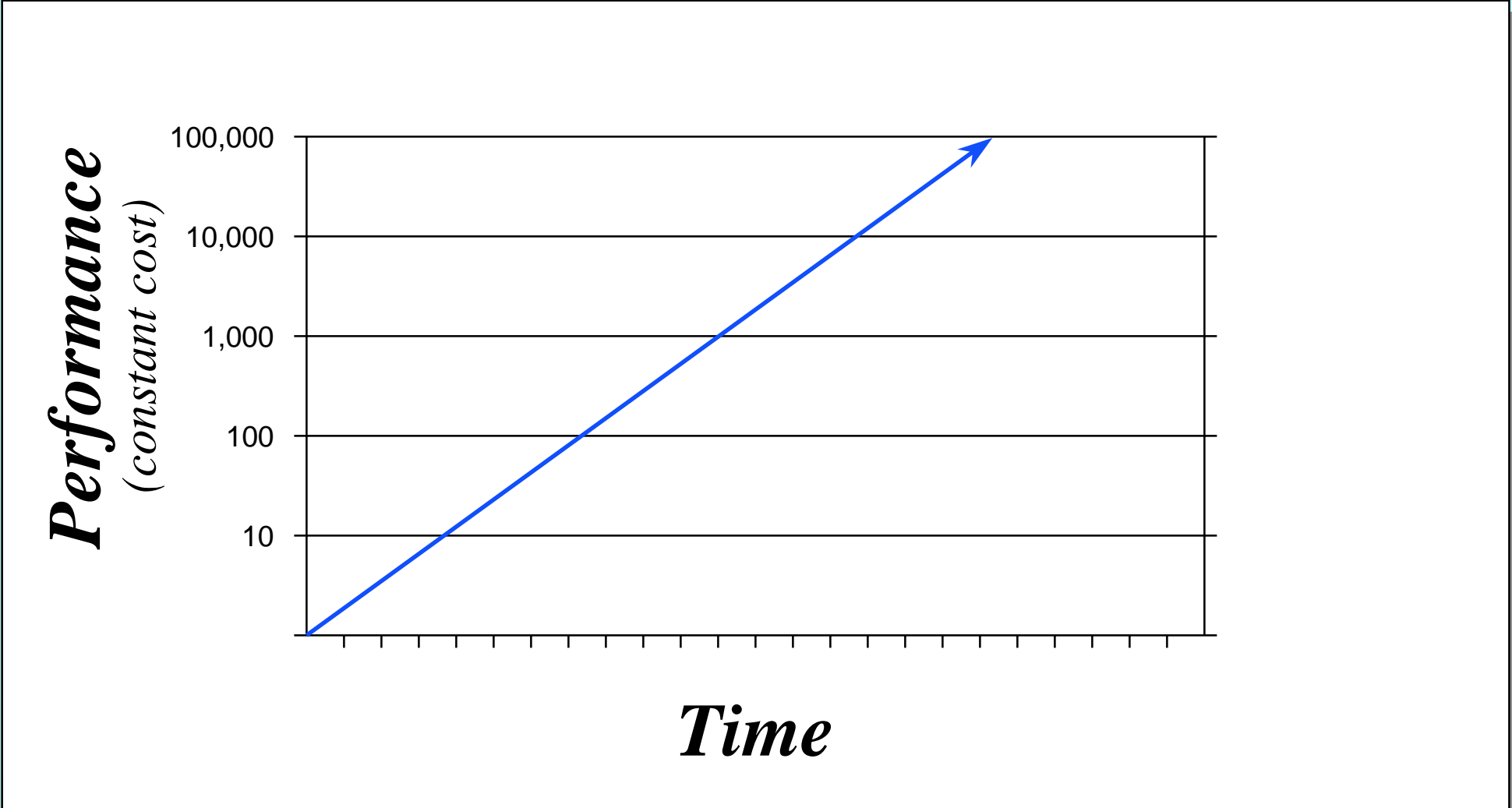
# Moore's Law: *The Statement*

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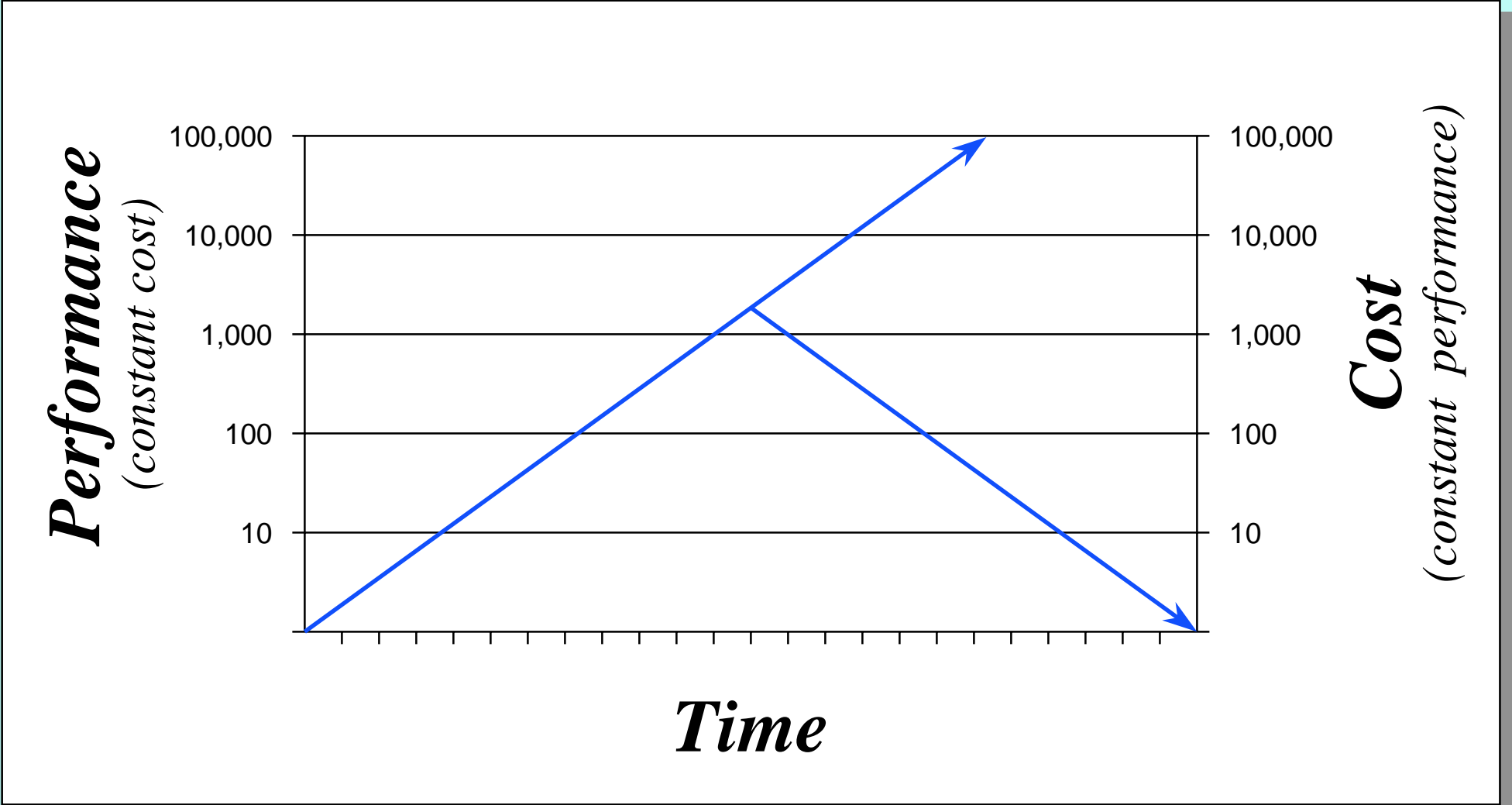
Every eighteen months, the number of transistors that can be placed on a chip doubles.

Gordon Moore, co-founder of Intel...

# Moore's Law: *The Effect*



# Moore's Law: *The Effect*



# **Moore's Law: *The Effect***

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## **Three Phases of Novel IT Applications**

- It's Impossible

# Moore's Law: *The Effect*

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## Three Phases of Novel IT Applications

- It's Impossible
- It's Impractical

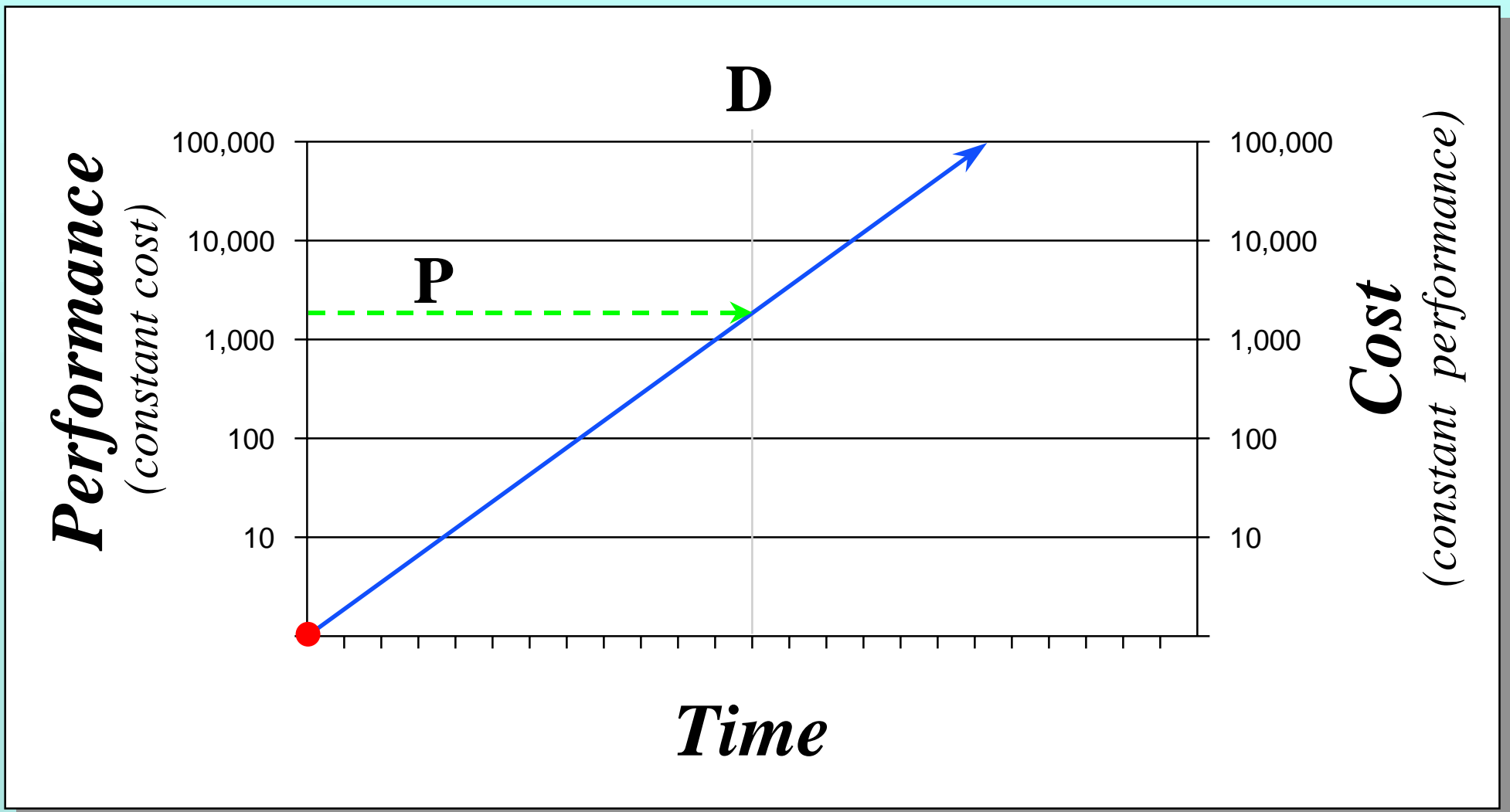
# Moore's Law: *The Effect*

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## Three Phases of Novel IT Applications

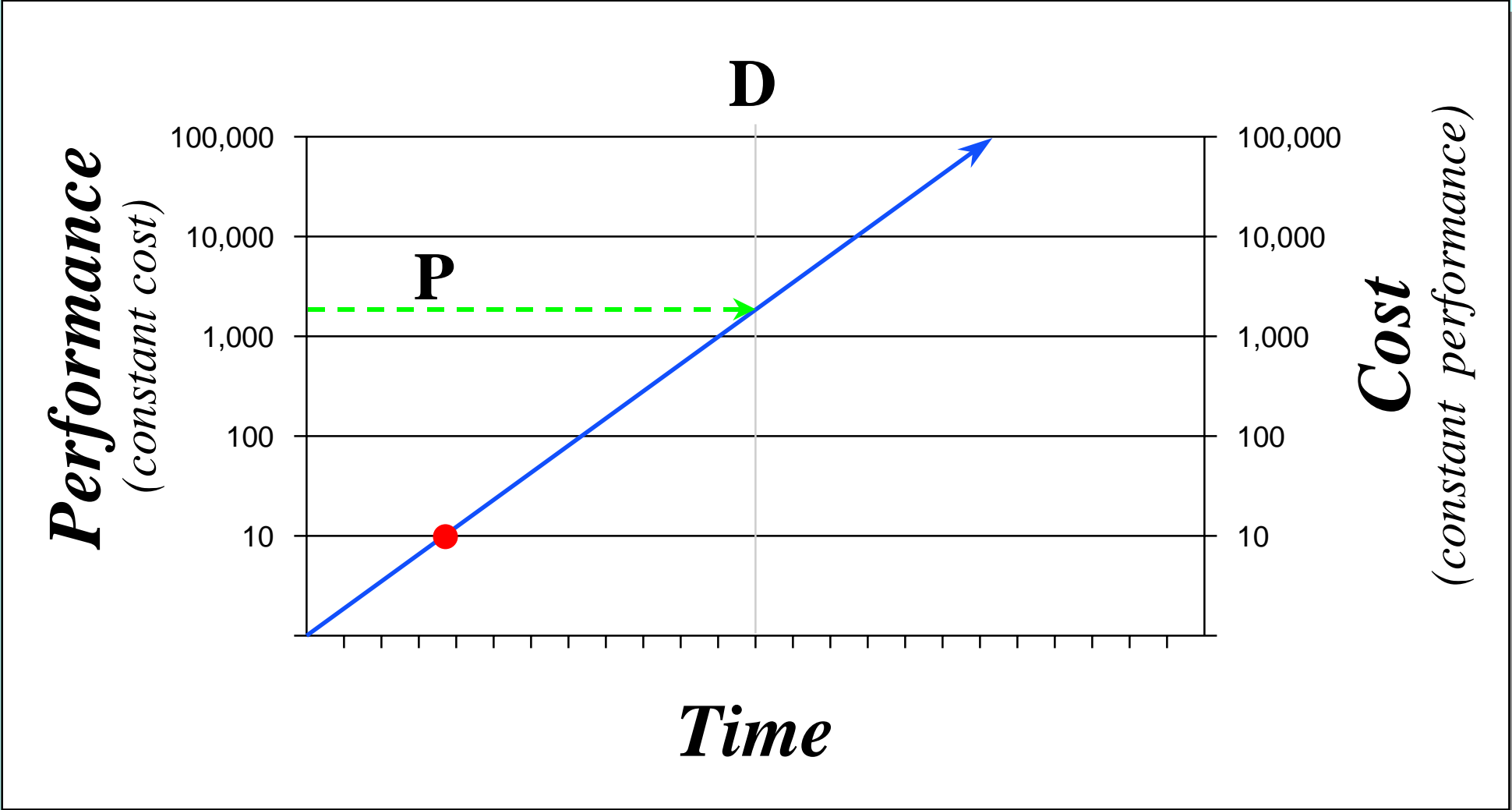
- It's Impossible
- It's Impractical
- It's Overdue

# Moore's Law: *The Effect*

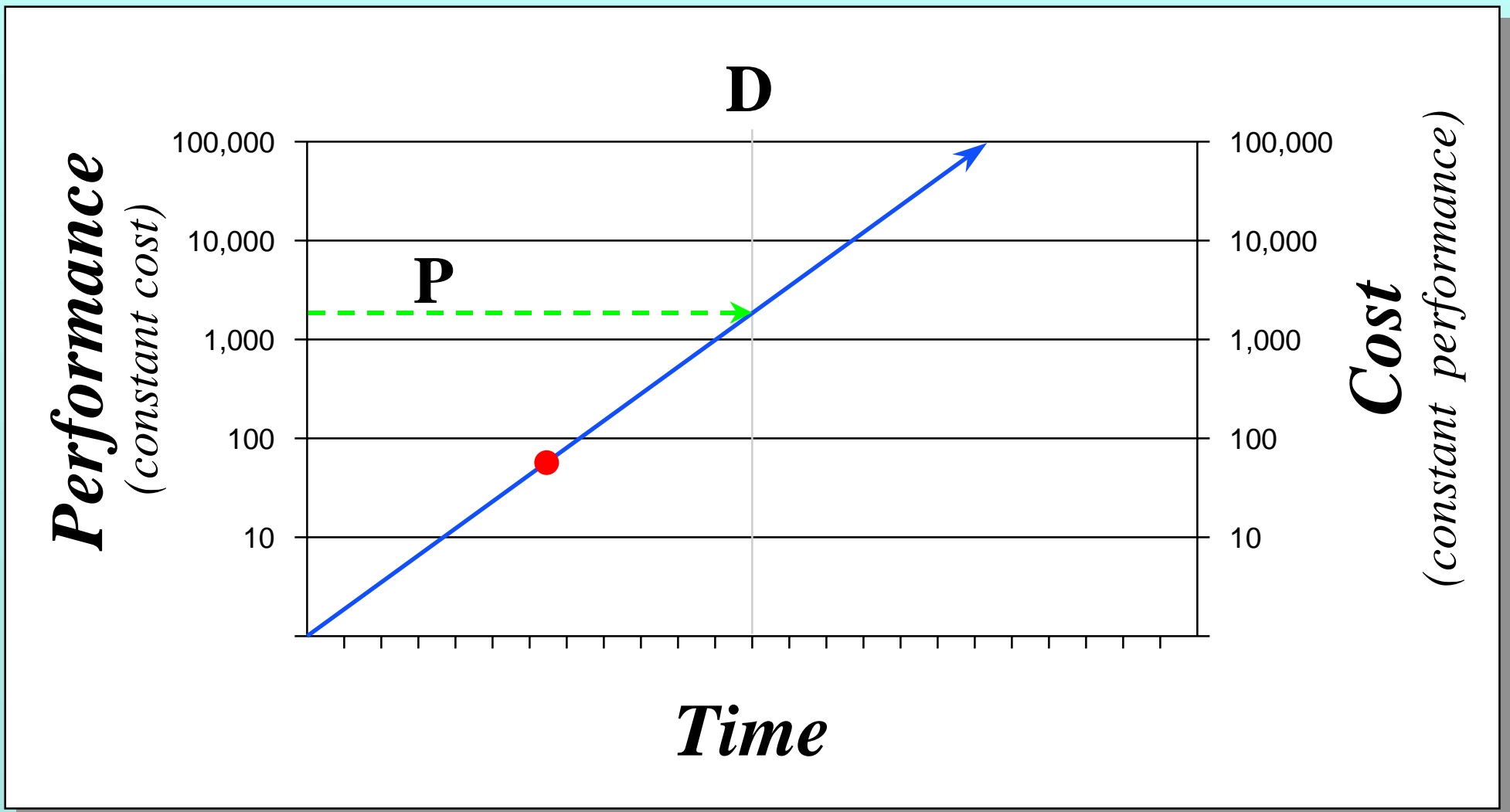




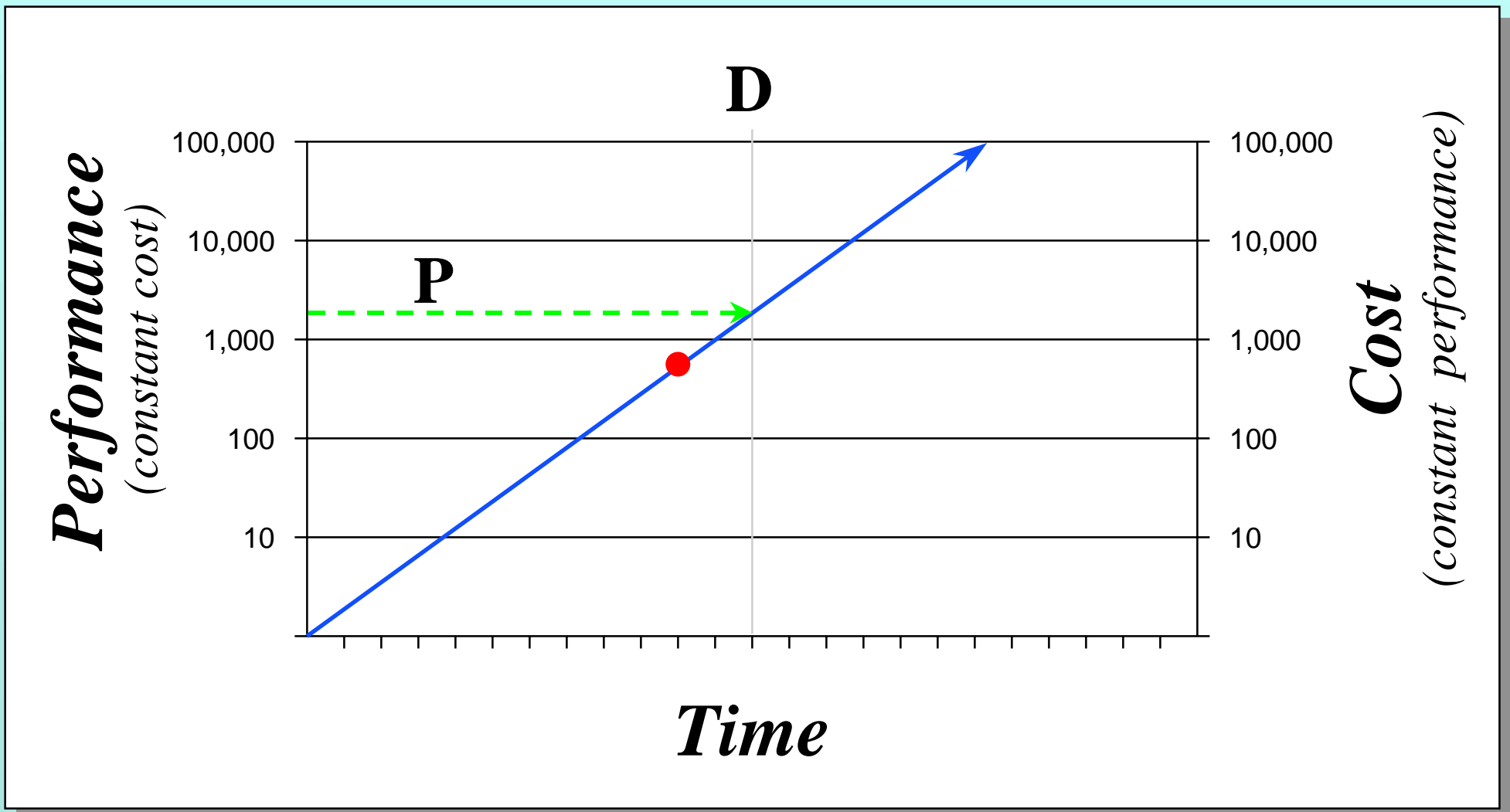
# Moore's Law: *The Effect*



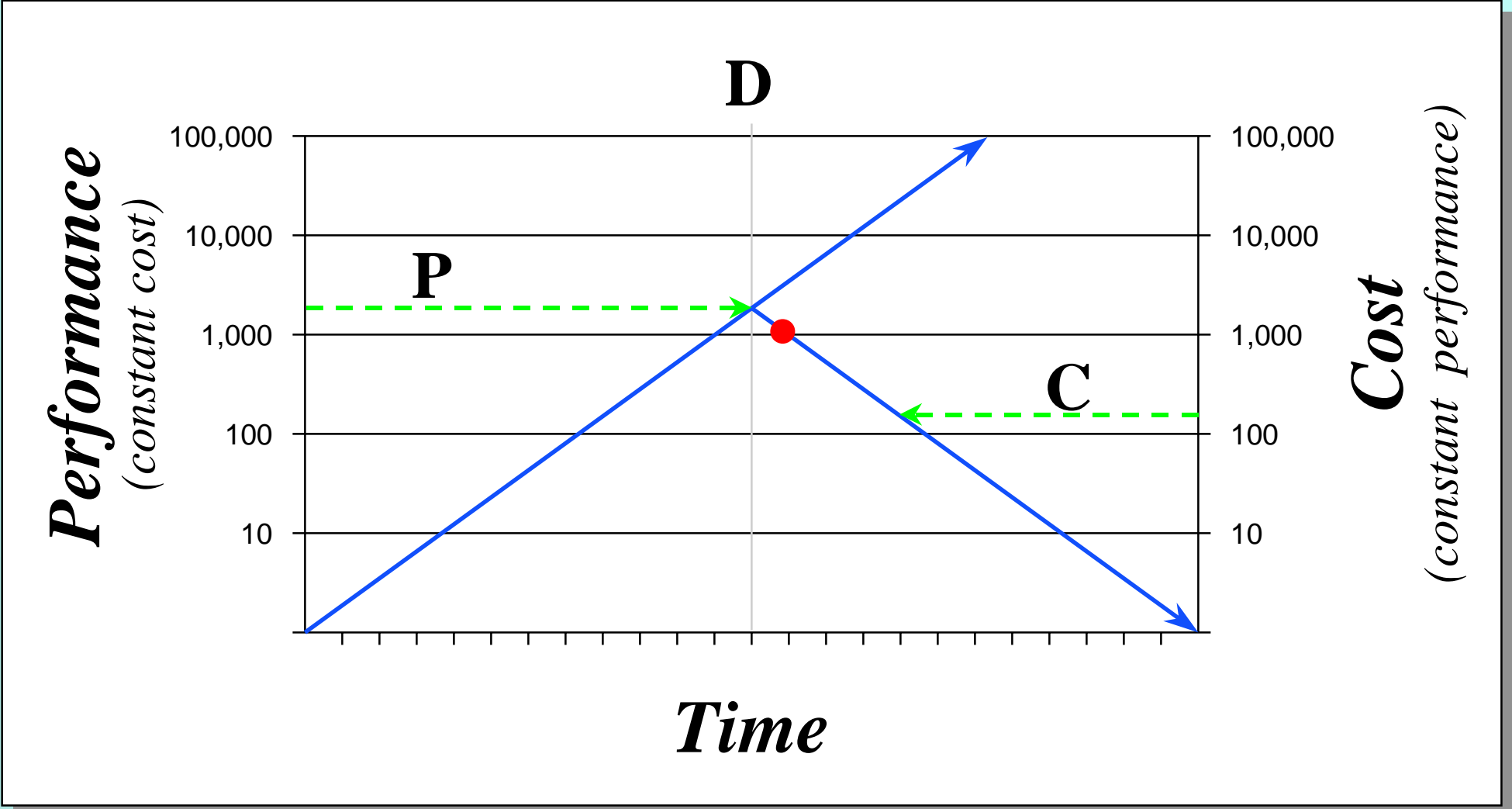
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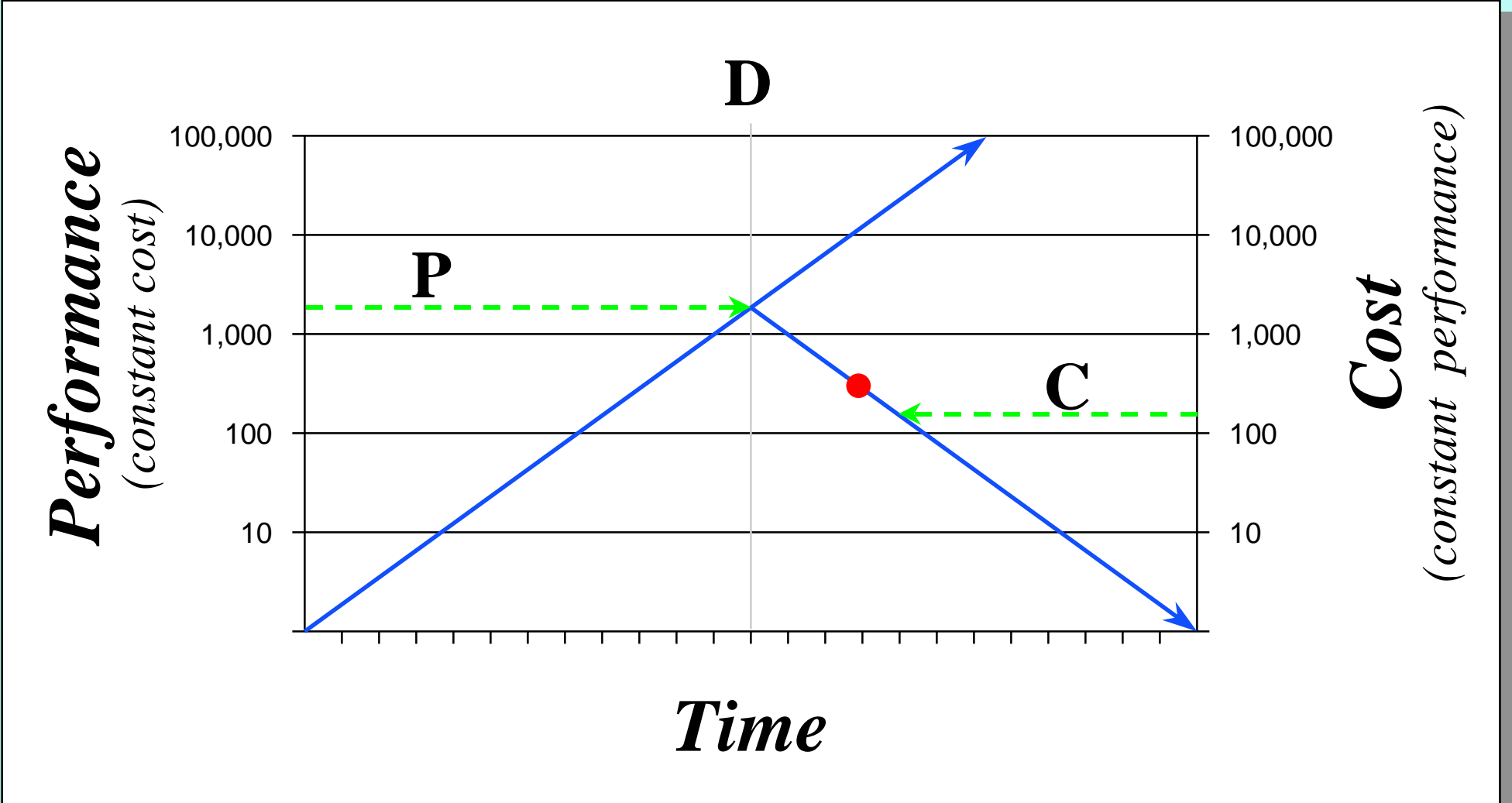
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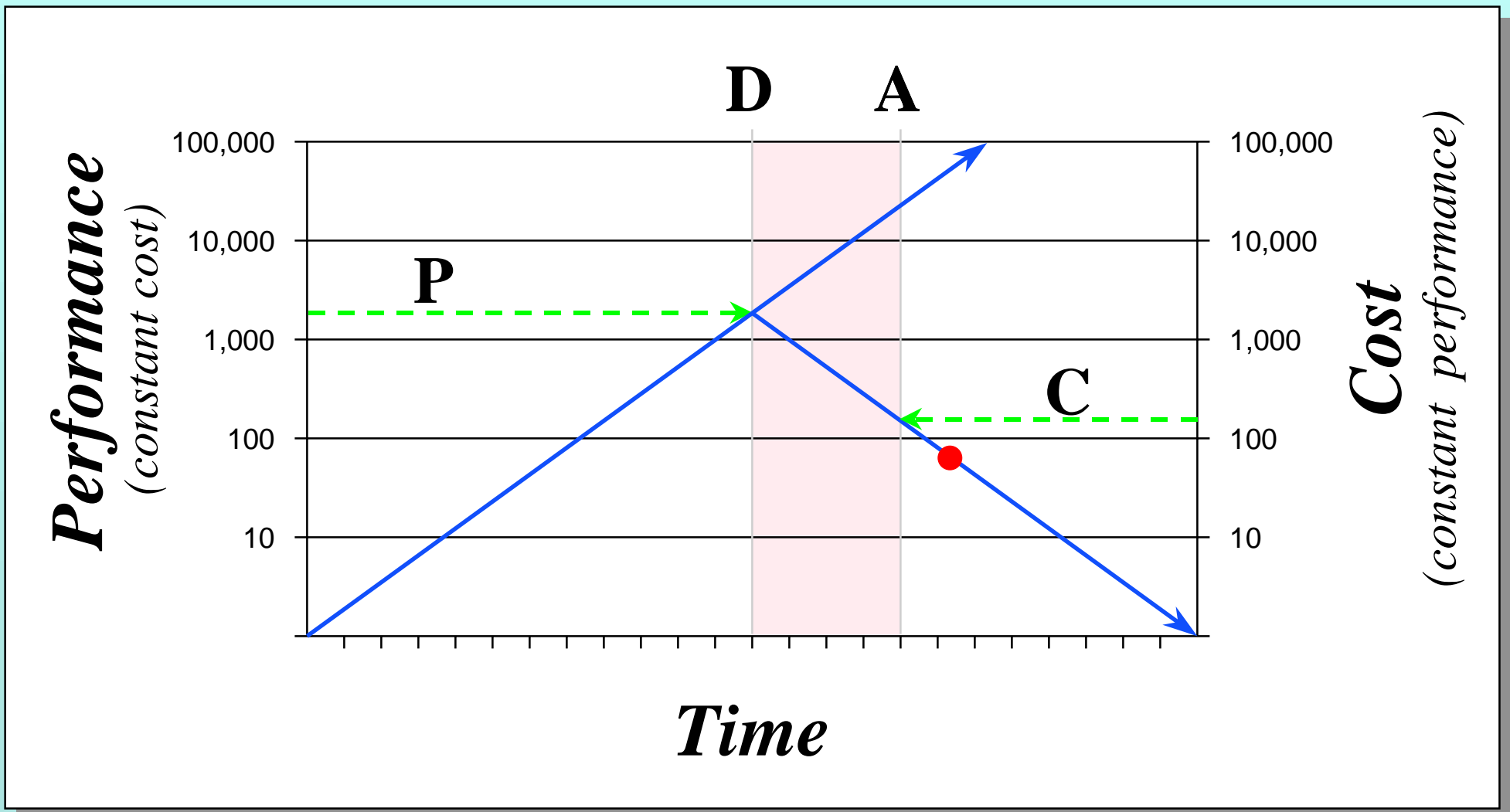
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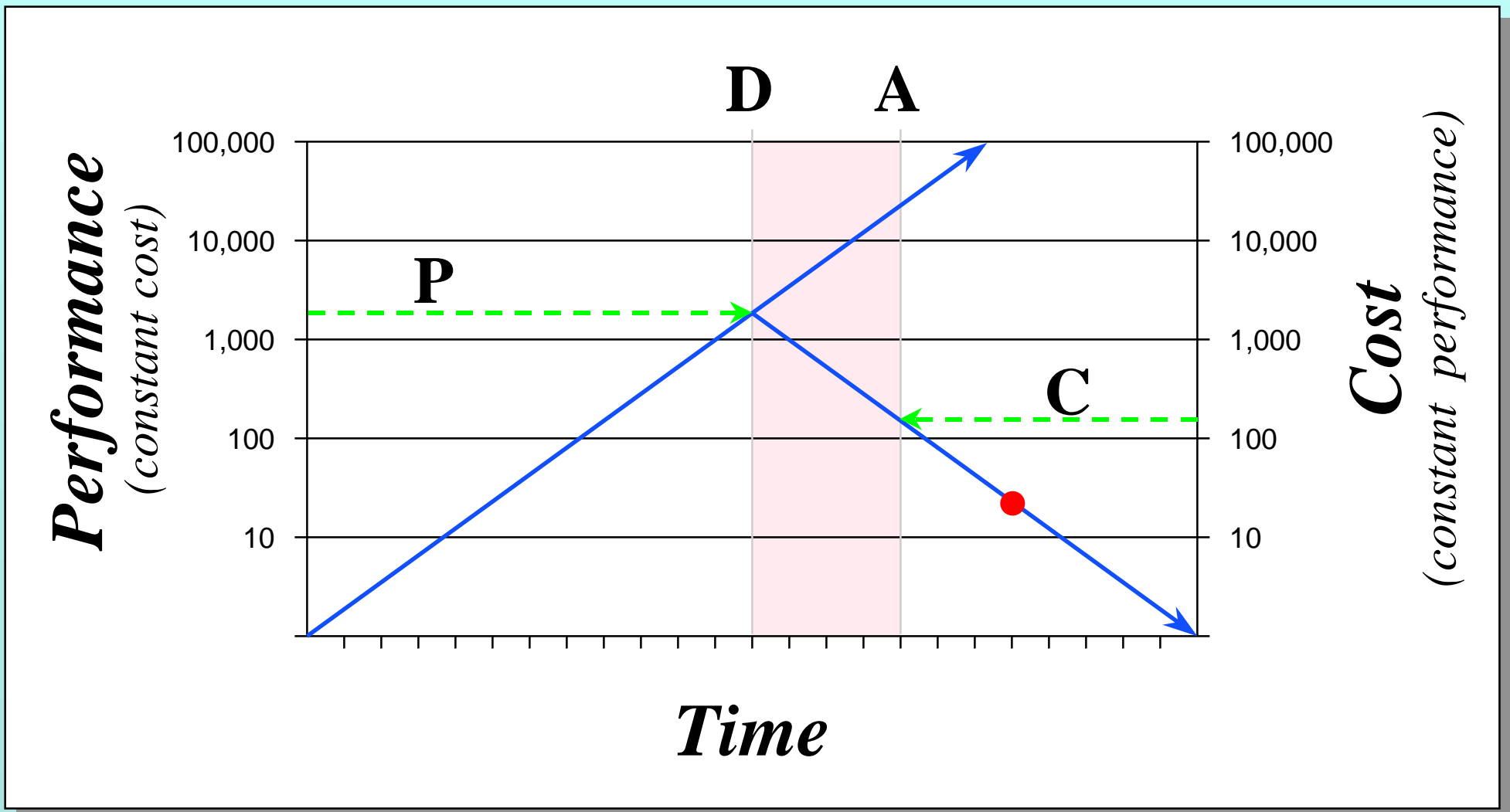
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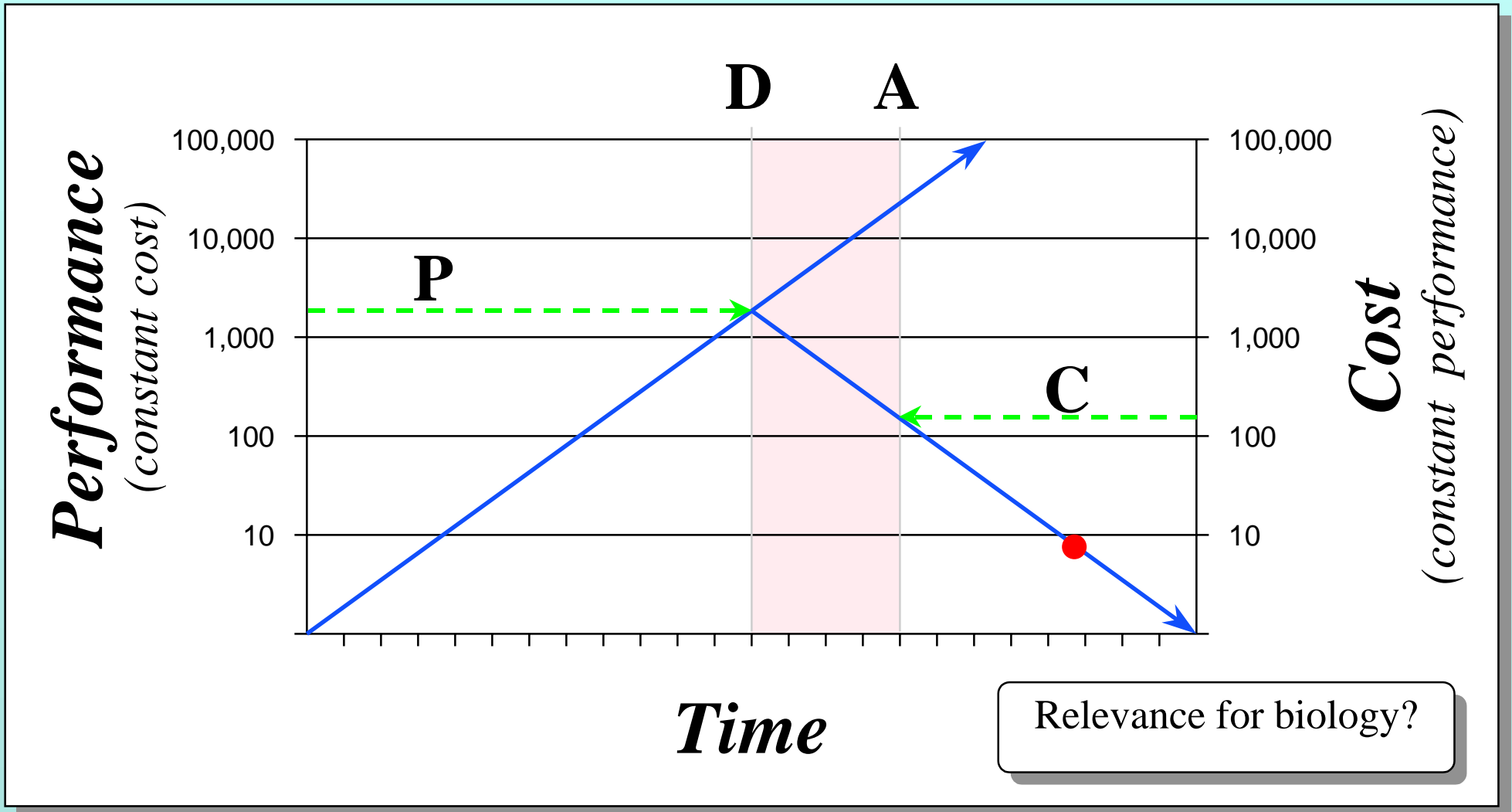
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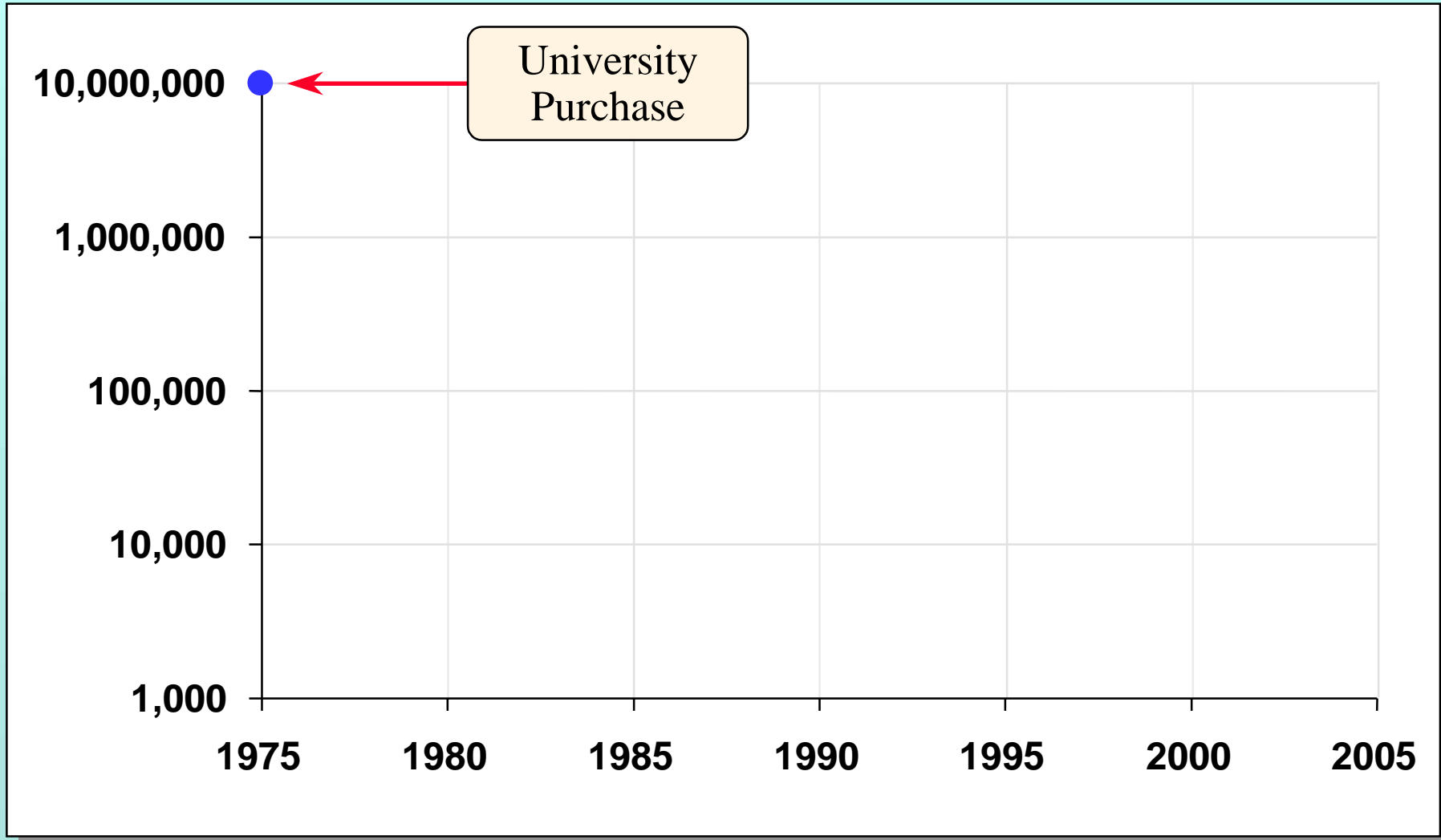


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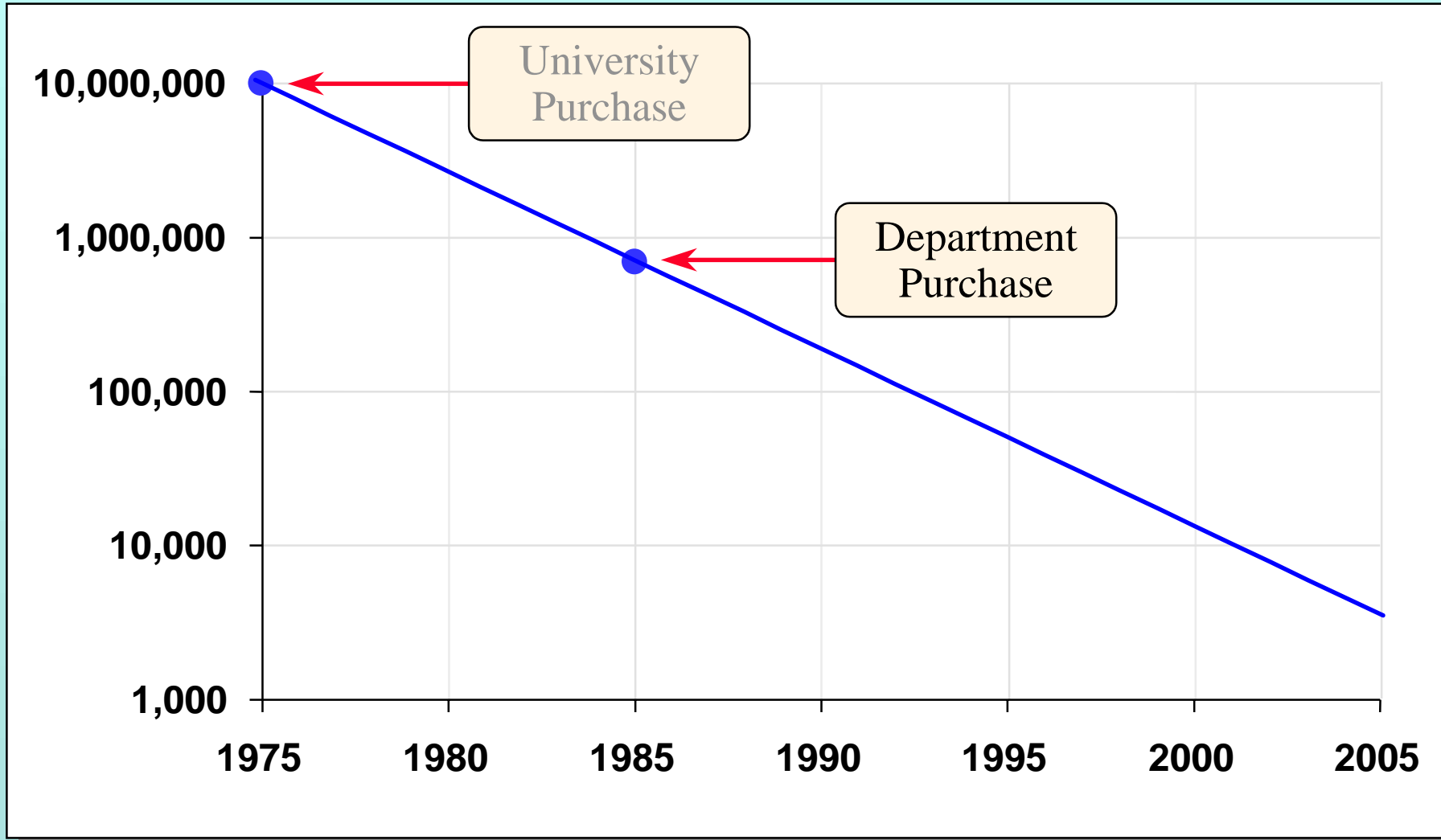




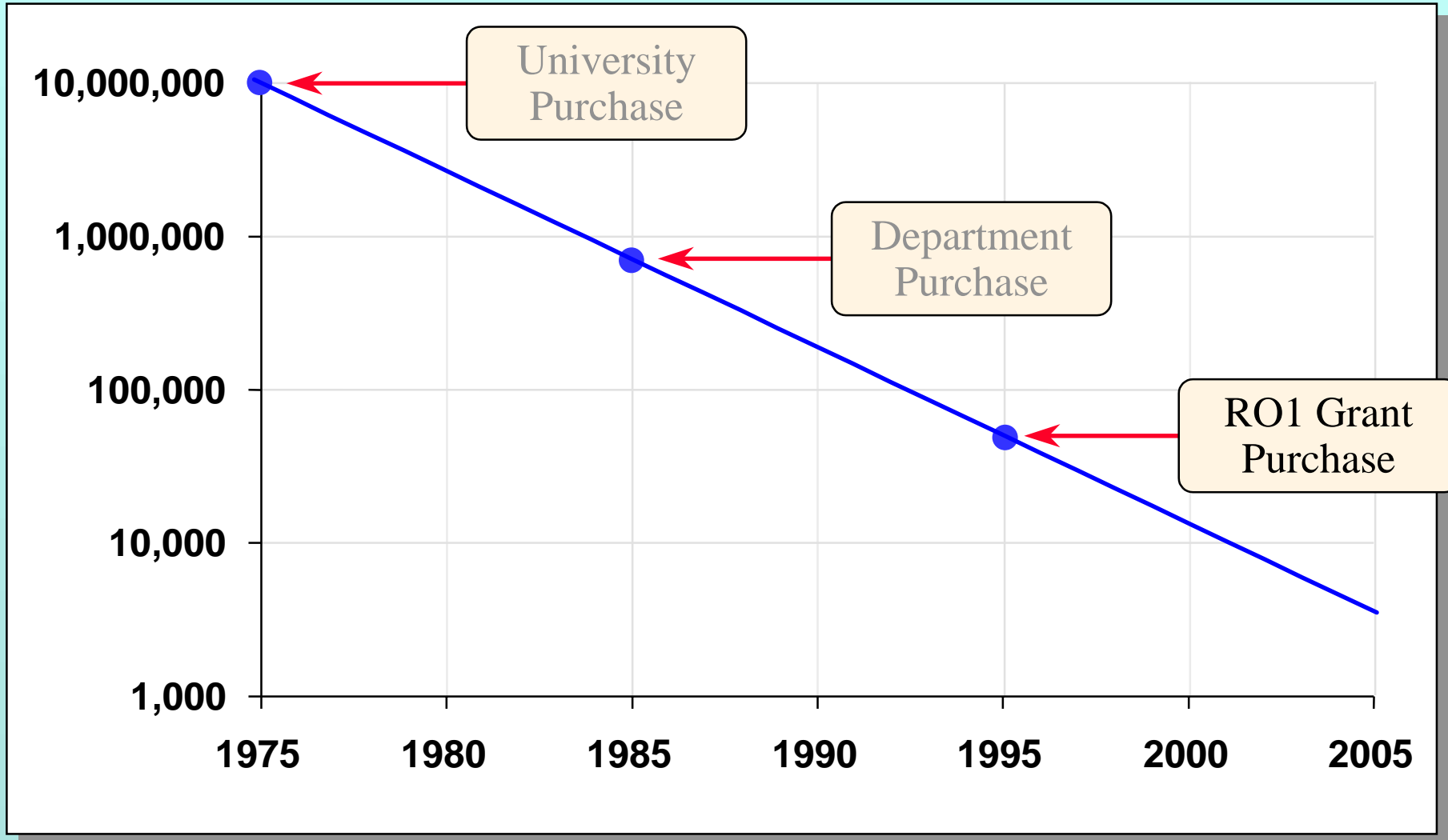
# Cost (constant performance)



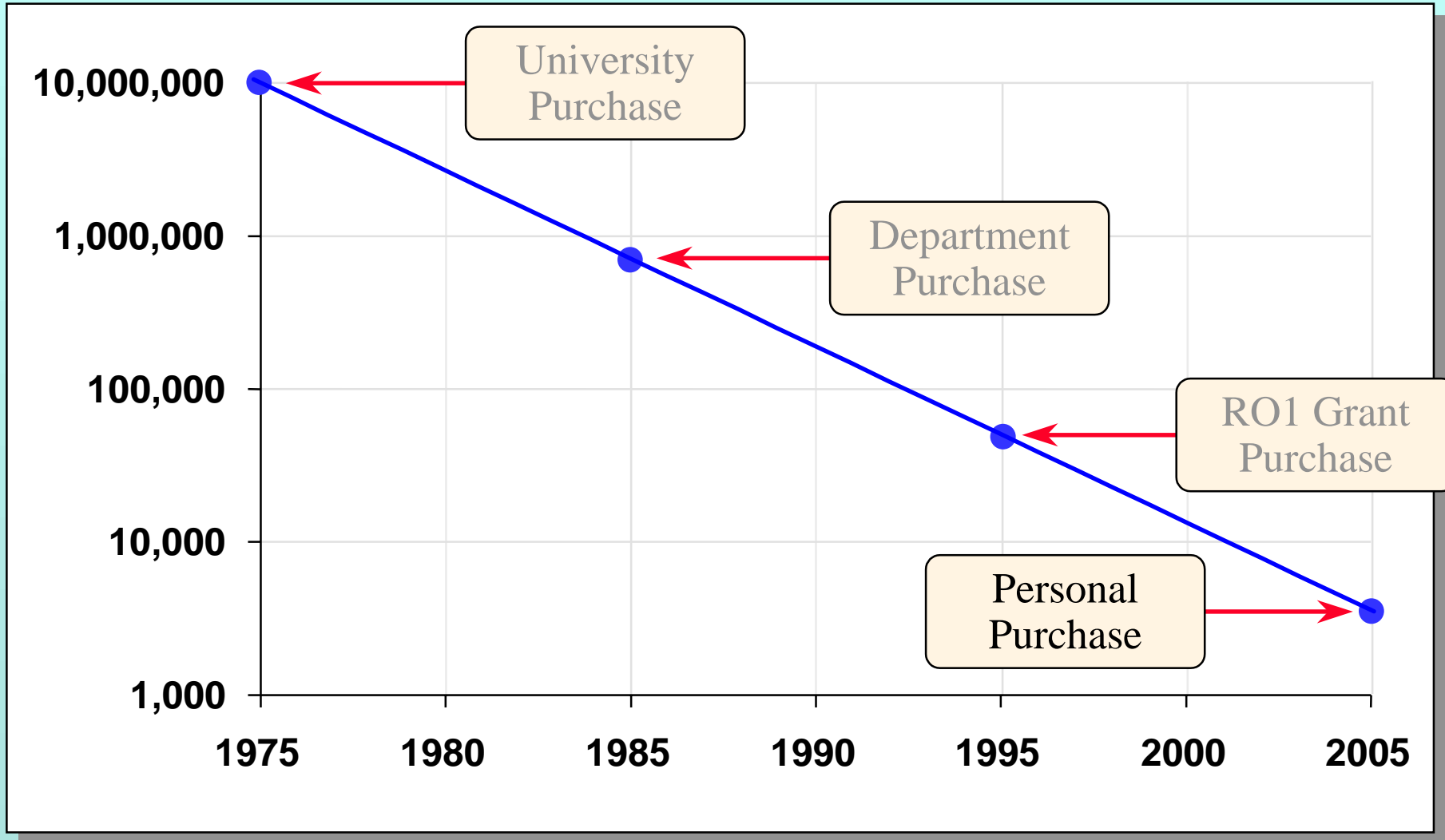
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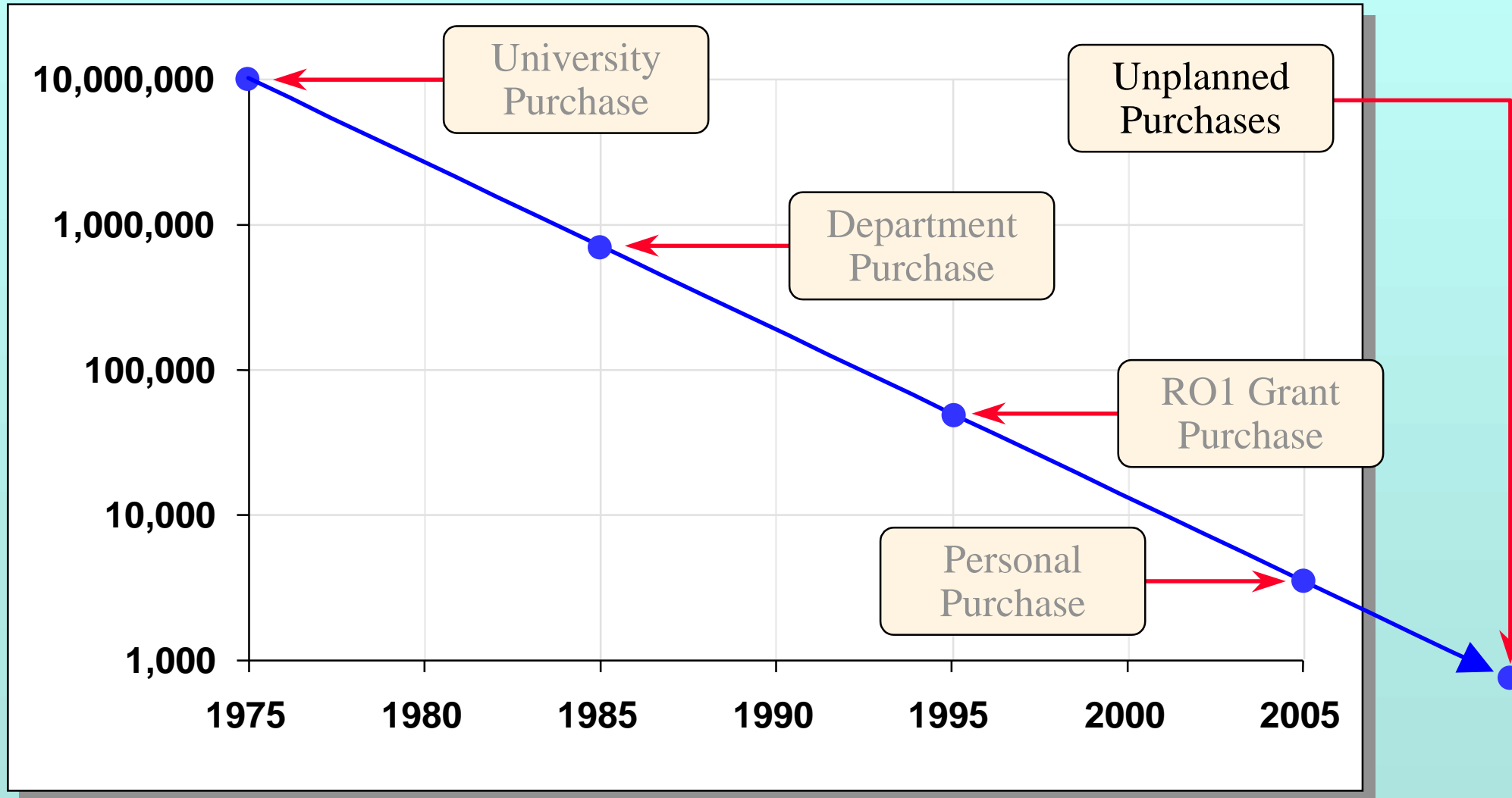
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# IT-Biology Synergism

# IT is Special

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Information Technology:

- *affects the performance **and** the management of tasks*

# IT is Special

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- *affects the performance and the management of tasks*
- *allows the manipulation of huge amounts of highly complex data*



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- *is incredibly plastic*  
*(programming and poetry are both exercises in pure thought)*

# IT is Special

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## Information Technology:

- *affects the performance and the management of tasks*
- *allows the manipulation of huge amounts of highly complex data*
- *is incredibly plastic*  
*(programming and poetry are both exercises in pure thought)*
- *improves exponentially* (Moore's Law)

# Biology is Special

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Life is Characterized by:

- *individuality*

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# Biology is Special

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No law of large numbers, since every living thing is genuinely unique.



# IT-Biology Synergism

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- *Physics needs calculus, the method for manipulating information about statistically large numbers of vanishingly small, independent, equivalent things.*

# IT-Biology Synergism

---

- *Physics needs calculus, the method for manipulating information about statistically large numbers of vanishingly small, independent, equivalent things.*
- *Biology needs information technology, the method for manipulating information about large numbers of dependent, historically contingent, individual things.*

# Biology is Special

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For it is in relation to the statistical point of view that the structure of the vital parts of living organisms differs so entirely from that of any piece of matter that we physicists and chemists have ever handled in our laboratories or mentally at our writing desks.

Erwin Schrödinger. 1944. *What is Life*.

# Genetics as Code

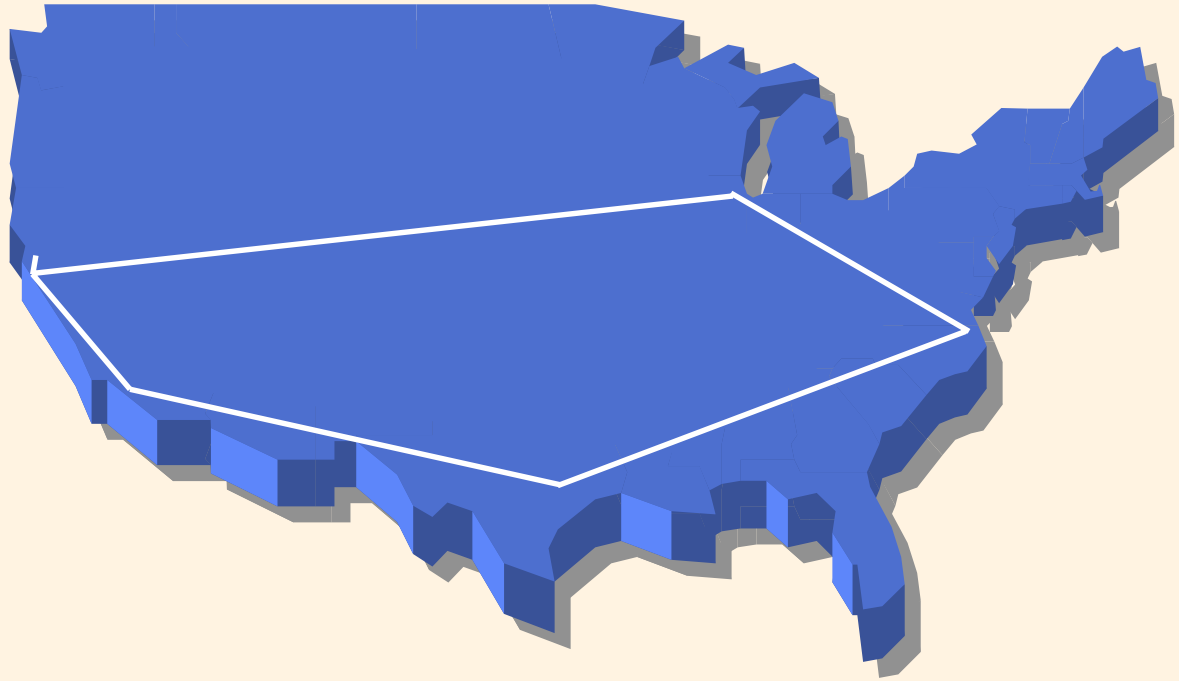
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[The] chromosomes ... contain in some kind of code-script the entire pattern of the individual's future development and of its functioning in the mature state. ... [By] code-script we mean that the all-penetrating mind, once conceived by Laplace, to which every causal connection lay immediately open, could tell from their structure whether [an egg carrying them] would develop, under suitable conditions, into a black cock or into a speckled hen, into a fly or a maize plant, a rhodo-dendron, a beetle, a mouse, or a woman.

Erwin Schrödinger. 1944. *What is Life*.

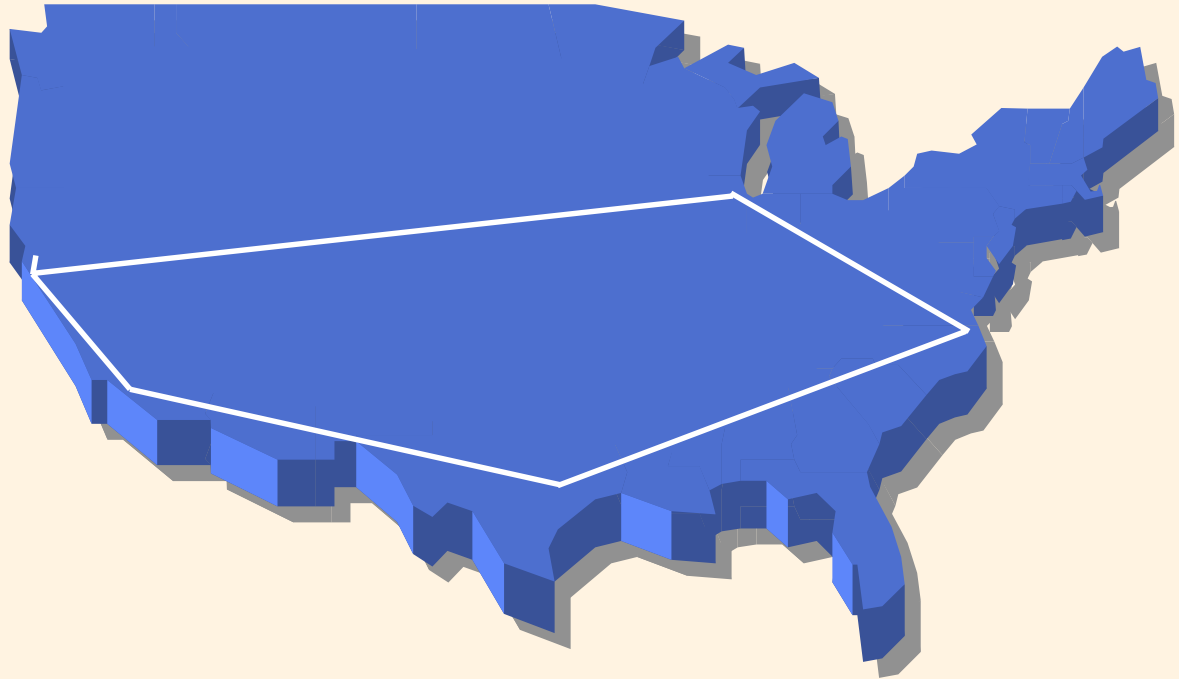
# One Human Sequence

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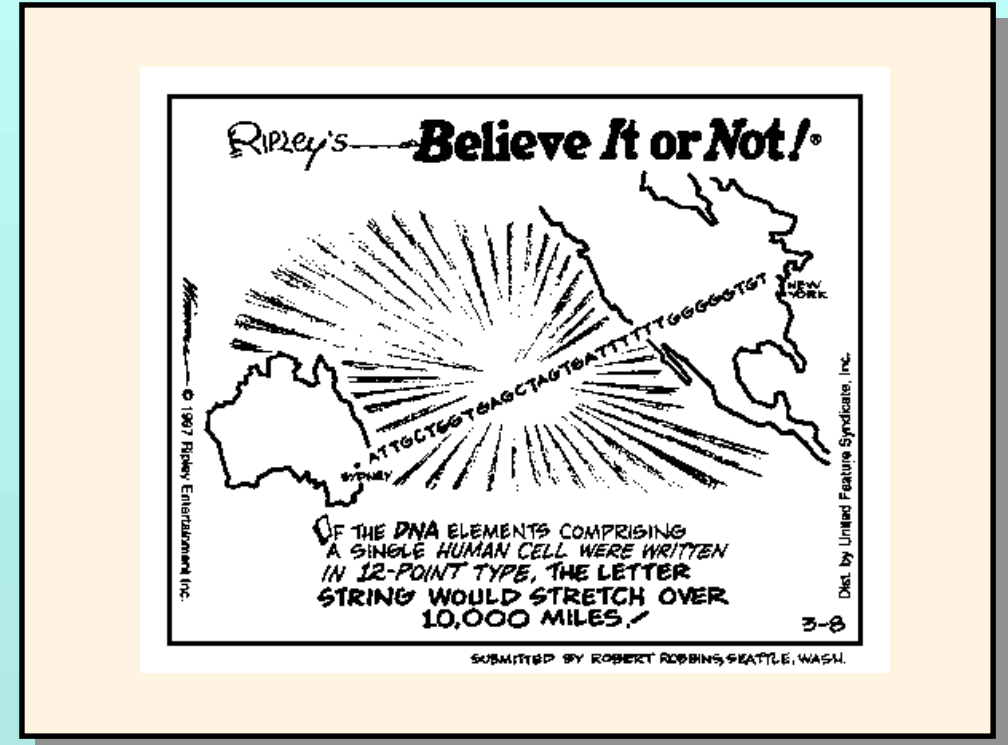
We now know that Schrödinger's mysterious human "code-script" consists of 3.3 billion base pairs of DNA.



Typed in 10-pitch font, one human sequence would stretch for more than 5,000 miles. Digitally formatted, it could be stored on one CD-ROM. Biologically encoded, it fits easily within a single cell.

# One Human Sequence

A variant of this factoid actually made it into *Ripley's Believe It or Not*, but that's another story...



For details on Ripley's interest in DNA, see

[HTTP://LX1.SELU.COM/~rjr/factoids/genlen.html](http://lx1.selu.com/~rjr/factoids/genlen.html)

# Bio-digital Information

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## **DNA is a highly efficient digital storage device:**

- There is more mass-storage capacity in the DNA of a side of beef than in all the hard drives of all the world's computers.



# Bio-digital Information

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- There is more mass-storage capacity in the DNA of a side of beef than in all the hard drives of all the world's computers.
- Storing all of the (redundant) information in all of the world's DNA on computer hard disks would require that the entire surface of the Earth be covered to a depth of three miles in Conner 1.0 gB drives.

# Genomics: An Example

# Human Genome Project - Goals

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- construction of a high-resolution genetic map of the human genome;

USDOE. 1990. *Understanding Our Genetic Inheritance. The U.S. Human Genome Project: The First Five Years.*

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- determination of the complete sequence of human DNA and of the DNA of selected model organisms;
- development of capabilities for collecting, storing, distributing, and analyzing the data produced;
- creation of appropriate technologies necessary to achieve these objectives.

USDOE. 1990. *Understanding Our Genetic Inheritance. The U.S. Human Genome Project: The First Five Years.*

# Infrastructure and the HGP

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Progress towards all of the [Genome Project] goals will require the establishment of well-funded centralized facilities, including a stock center for the cloned DNA fragments generated in the mapping and sequencing effort and a data center for the computer-based collection and distribution of large amounts of DNA sequence information.

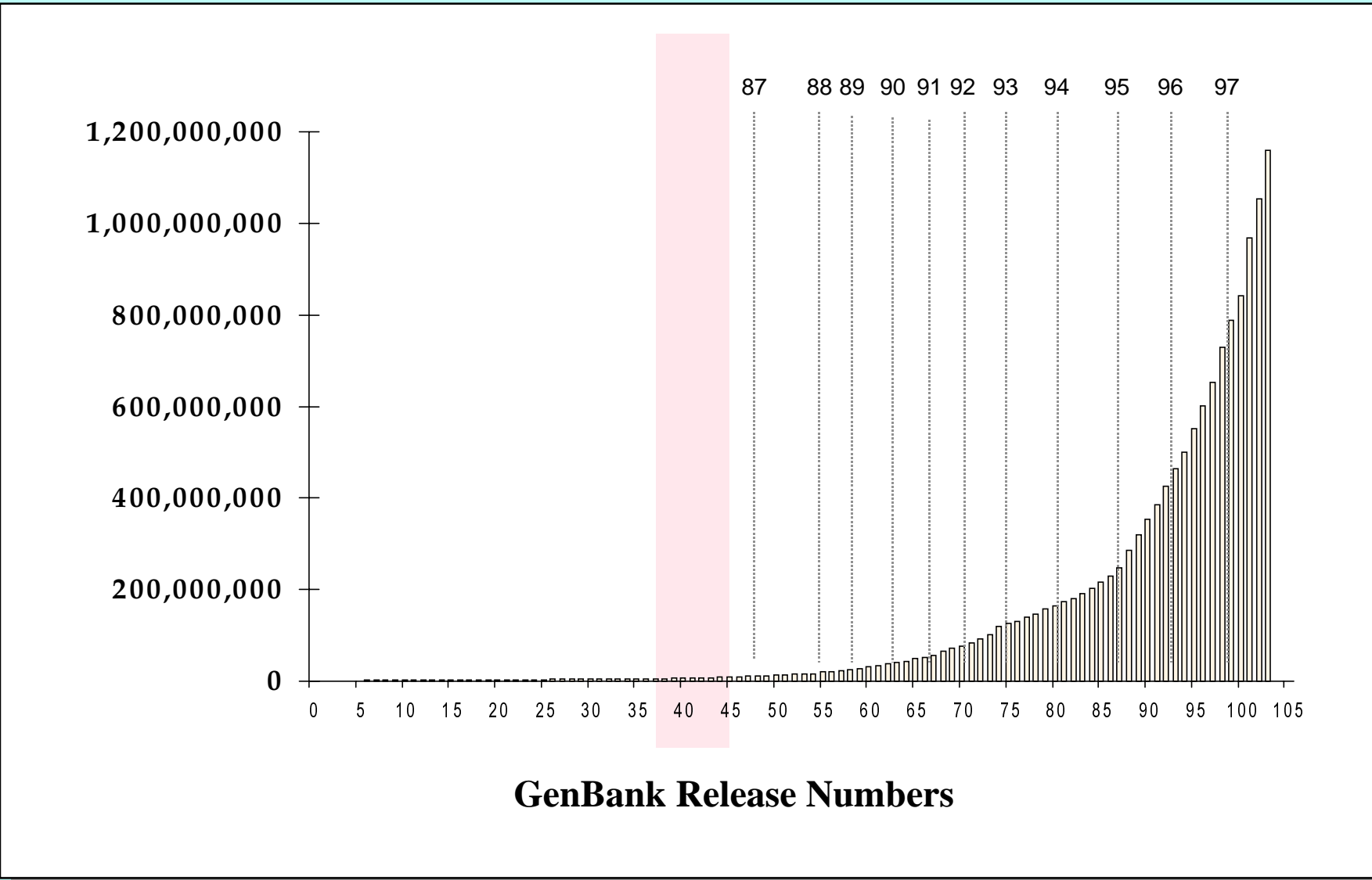
National Research Council. 1988. *Mapping and Sequencing the Human Genome*. Washington, DC: National Academy Press. p. 3



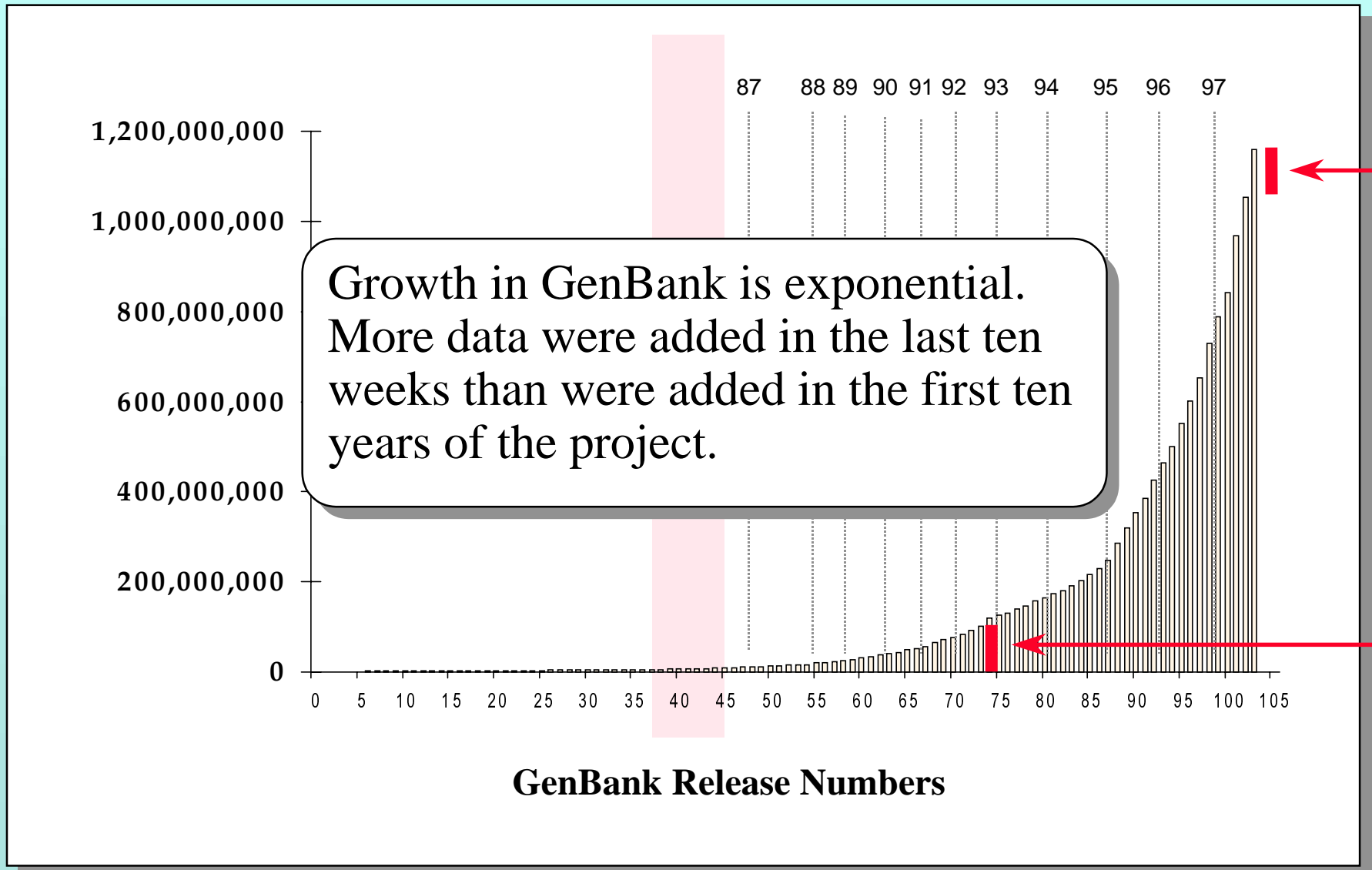
# GenBank Totals *(Release 103)*

| DIVISION                                 | Entries          | Per Cent        | Base Pairs           | Per Cent        |
|--|------------------|-----------------|----------------------|-----------------|
| Phage Sequences (PHG)                    | 1,313            | 0.074%          | 2,138,810            | 0.184%          |
| Viral Sequences (VRL)                    | 45,355           | 2.568%          | 44,484,848           | 3.834%          |
| Bacteria (BCT)                           | 38,023           | 2.153%          | 88,576,641           | 7.634%          |
| Plant, Fungal, and Algal Sequences (PLN) | 44,553           | 2.523%          | 92,259,434           | 7.951%          |
| Invertebrate Sequences (INV)             | 29,657           | 1.679%          | 105,703,550          | 9.110%          |
| Rodent Sequences (ROD)                   | 36,967           | 2.093%          | 45,437,309           | 3.916%          |
| Primate Sequences (PRI1-2)               | 75,587           | 4.280%          | 134,944,314          | 11.630%         |
| Other Mammals (MAM)                      | 12,744           | 0.722%          | 12,358,310           | 1.065%          |
| Other Vertebrate Sequences (VRT)         | 17,713           | 1.003%          | 17,040,159           | 1.469%          |
| High-Throughput Genome Sequences (HTG)   | 1,120            | 0.063%          | 72,064,395           | 6.211%          |
| Genome Survey Sequences (GSS)            | 42,628           | 2.414%          | 22,783,326           | 1.964%          |
| Structural RNA Sequences (RNA)           | 4,802            | 0.272%          | 2,487,397            | 0.214%          |
| Sequence Tagged Sites Sequences (STS)    | 52,824           | 2.991%          | 18,161,532           | 1.565%          |
| Patent Sequences (PAT)                   | 87,767           | 4.970%          | 27,593,724           | 2.378%          |
| Synthetic Sequences (SYN)                | 2,577            | 0.146%          | 5,698,945            | 0.491%          |
| Unannotated Sequences (UNA)              | 2,480            | 0.140%          | 1,933,676            | 0.167%          |
| EST1-17                                  | <b>1,269,737</b> | <b>71.905%</b>  | <b>466,634,317</b>   | <b>40.217%</b>  |
| TOTALS                                   | <b>1,765,847</b> | <b>100.000%</b> | <b>1,160,300,687</b> | <b>100.000%</b> |

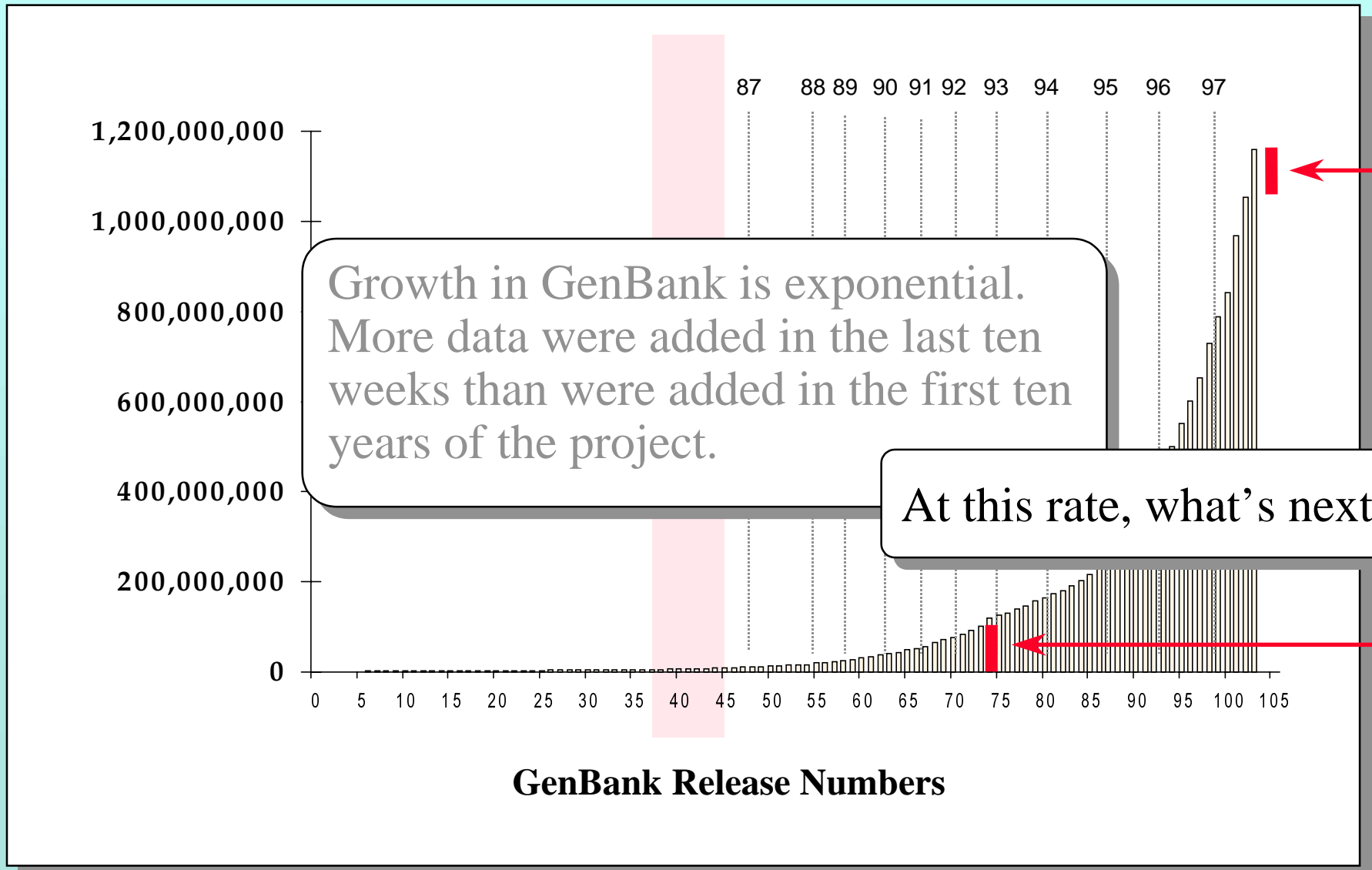
# Base Pairs in GenBank



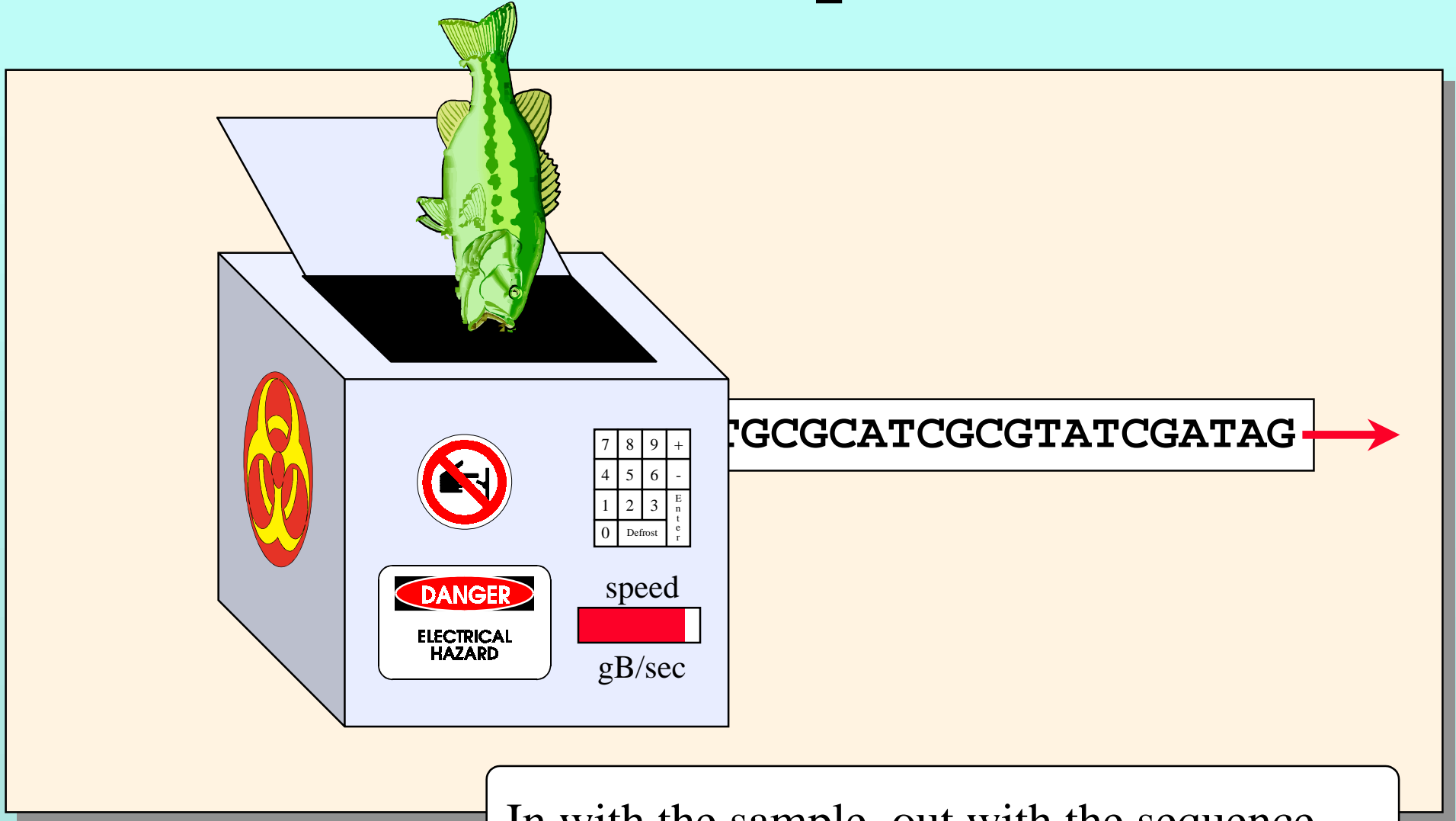
# Base Pairs in GenBank



# Base Pairs in GenBank



# ABI Bass-o-Matic Sequencer



In with the sample, out with the sequence...

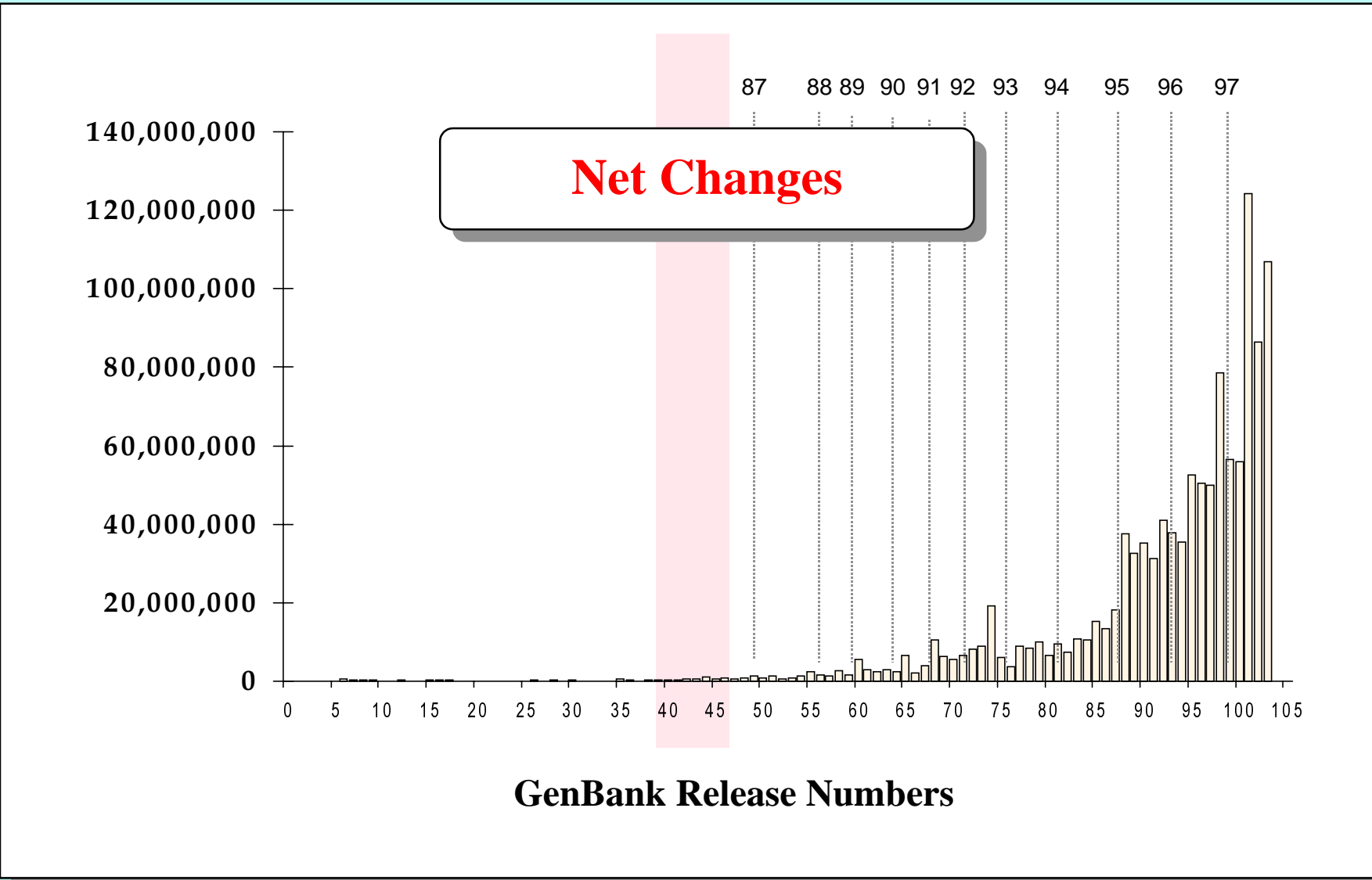
# What's Really Next

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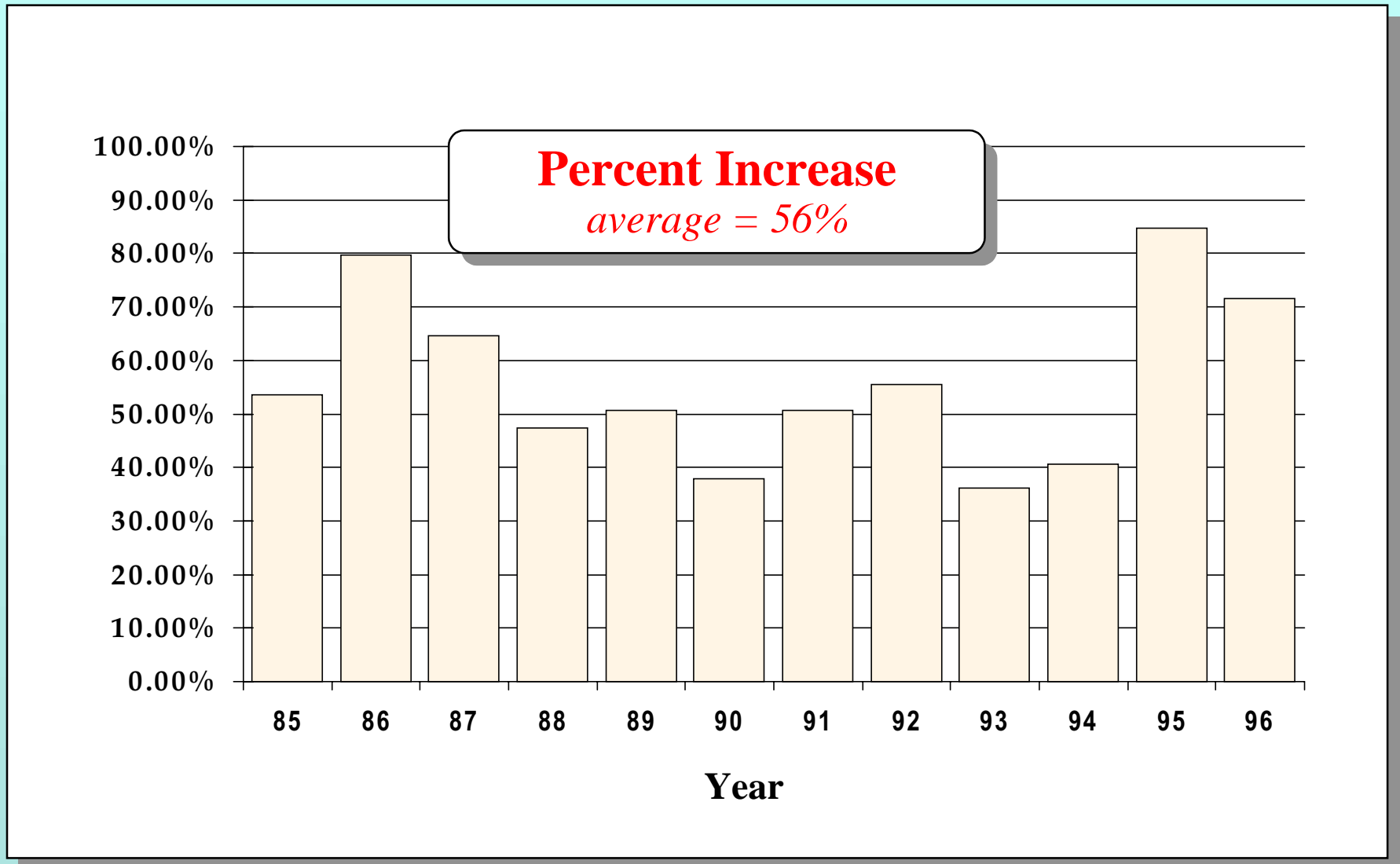
The post-genome era in biological research will take for granted ready access to huge amounts of genomic data.

The challenge will be *understanding* those data and using the understanding to solve real-world problems...

# Base Pairs in GenBank

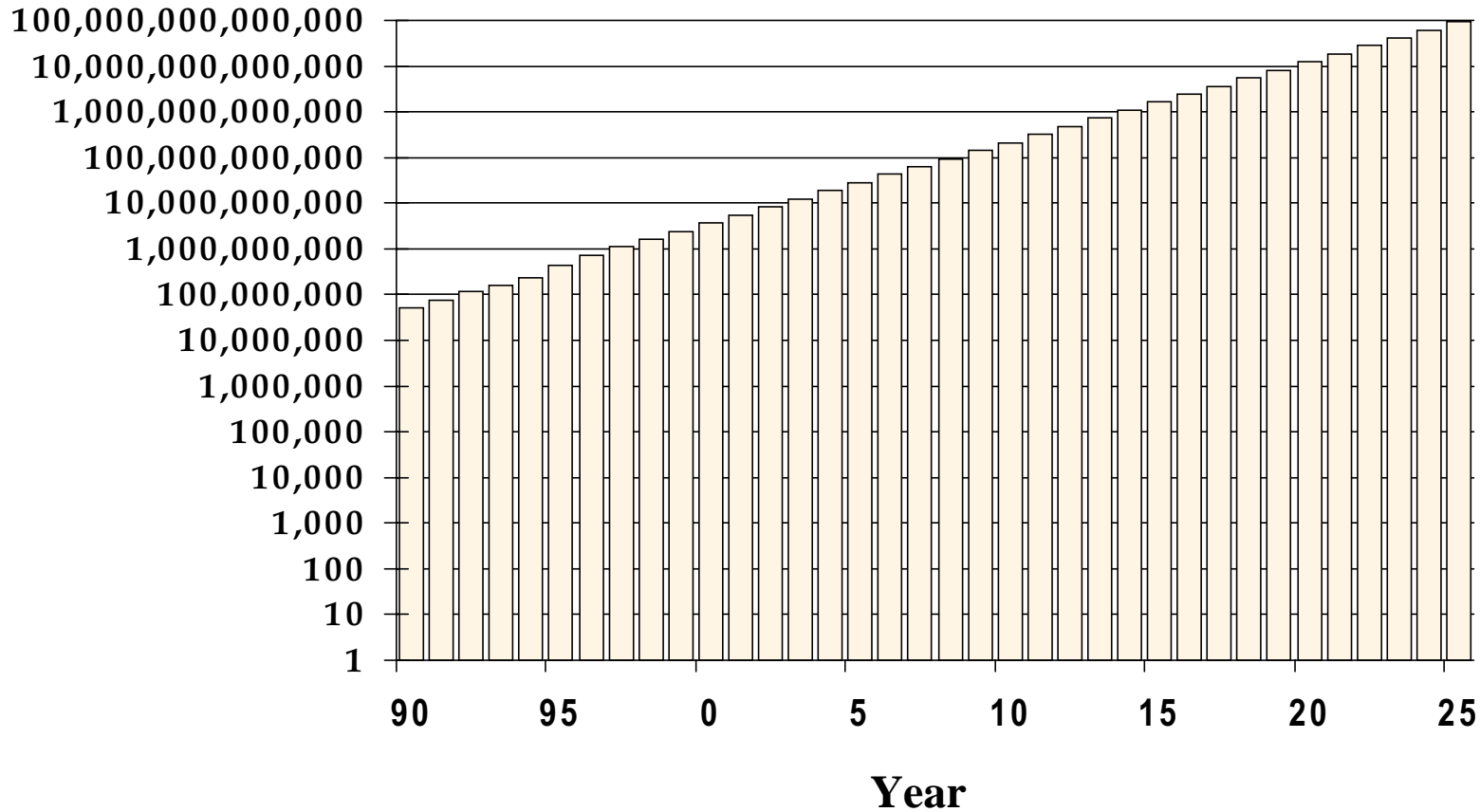


# Base Pairs in GenBank (*Percent Increase*)



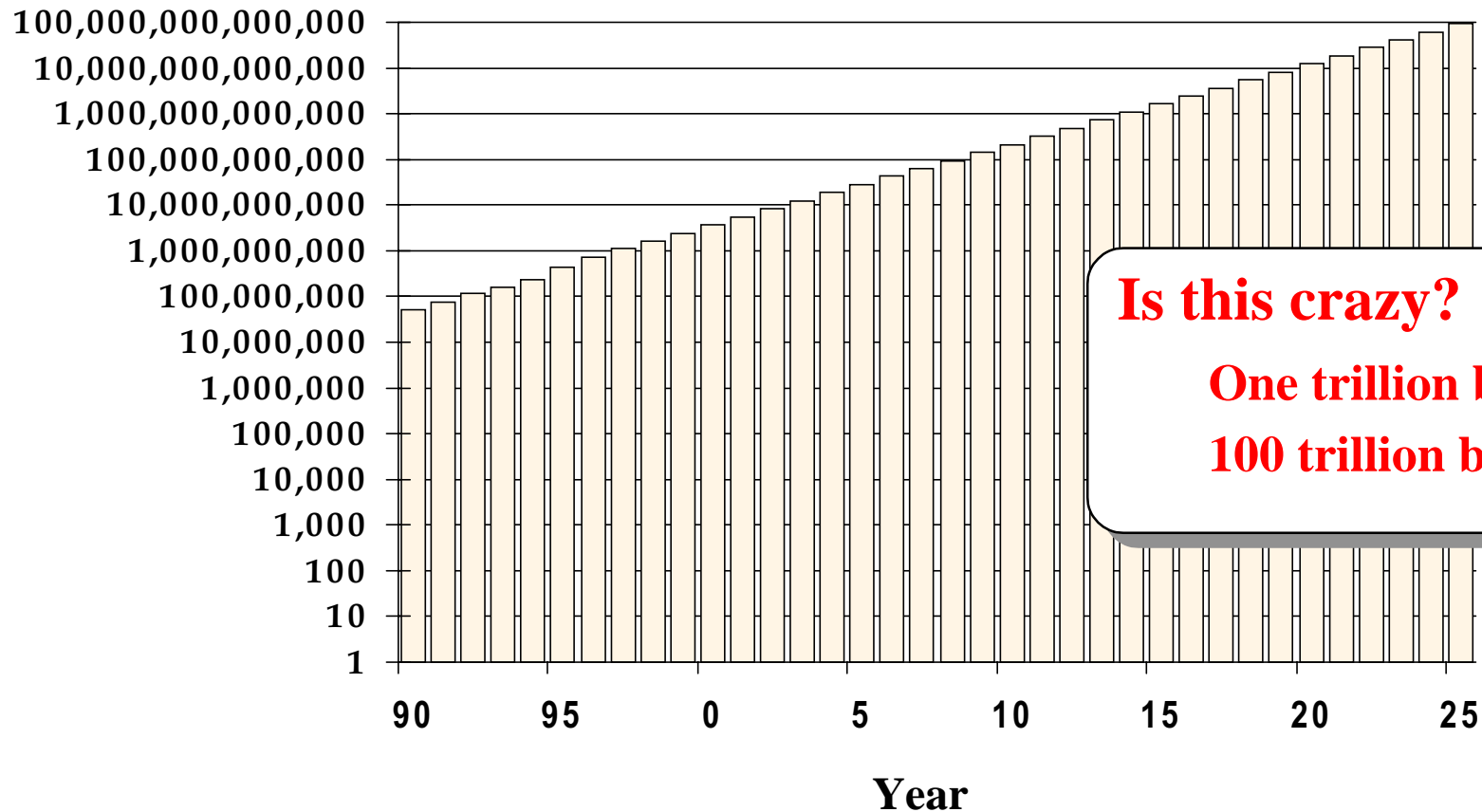


# Projected Base Pairs



Assumed annual growth rate: 50%  
*(less than current rate)*

# Projected Base Pairs



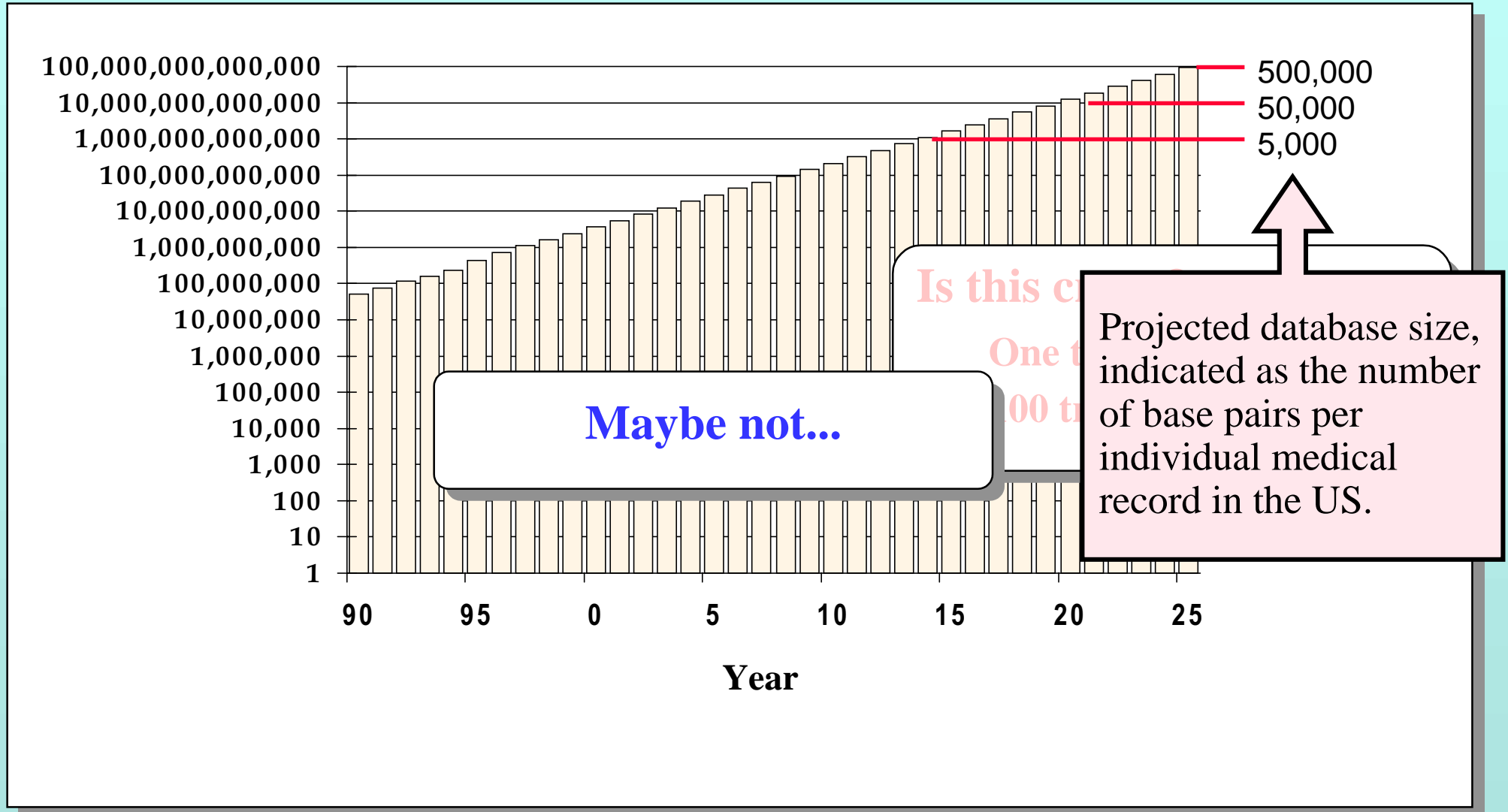
**Is this crazy?**

**One trillion bp by 2015**

**100 trillion by 2025**

Assumed annual growth rate: 50%  
*(less than current rate)*

# Projected Base Pairs



# 21st Century Biology

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*Post-Genome Era*

# The Post-Genome Era

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## **Post-genome research involves:**

- applying genomic tools and knowledge to more general problems
- asking new questions, tractable only to genomic or post-genomic analysis
- moving beyond the structural genomics of the human genome project and into the functional genomics of the post-genome era

# The Post-Genome Era

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## Suggested definition:

- functional genomics = biology

# The Post-Genome Era

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## An early analysis:

Walter Gilbert. 1991. Towards a paradigm shift in biology. *Nature*, 349:99.

# Paradigm Shift in Biology

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To use [the] flood of knowledge, which will pour across the computer networks of the world, biologists not only must become computer literate, but also change their approach to the problem of understanding life.

Walter Gilbert. 1991. Towards a paradigm shift in biology. *Nature*, 349:99.



# Paradigm Shift in Biology

---

The new paradigm, now emerging, is that all the 'genes' will be known (in the sense of being resident in databases available electronically), and that the starting point of a biological investigation will be theoretical. An individual scientist will begin with a theoretical conjecture, only then turning to experiment to follow or test that hypothesis.

Walter Gilbert. 1991. Towards a paradigm shift in biology. *Nature*, 349:99.

# Paradigm Shift in Biology

## Case of Microbiology

< 5,000 known and described bacteria

5,000,000 base pairs per genome

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25,000,000,000 TOTAL base pairs

If a full, annotated sequence were available for all known bacteria, the practice of microbiology would match Gilbert's prediction.

# 21st Century Biology

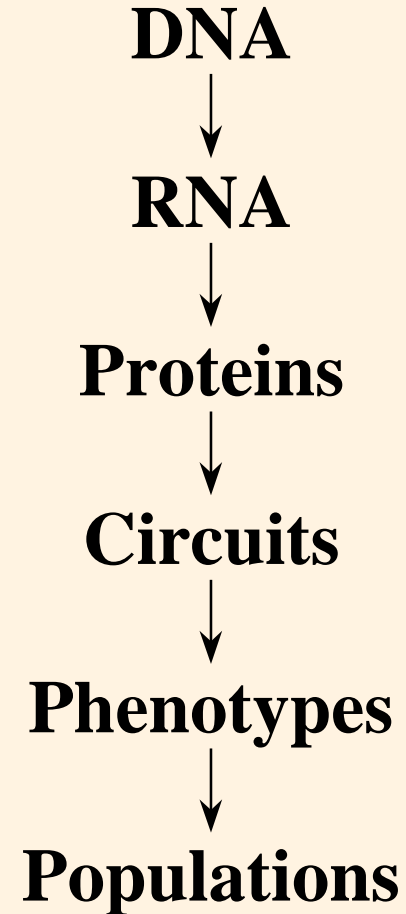
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*The Science*

# Fundamental Dogma

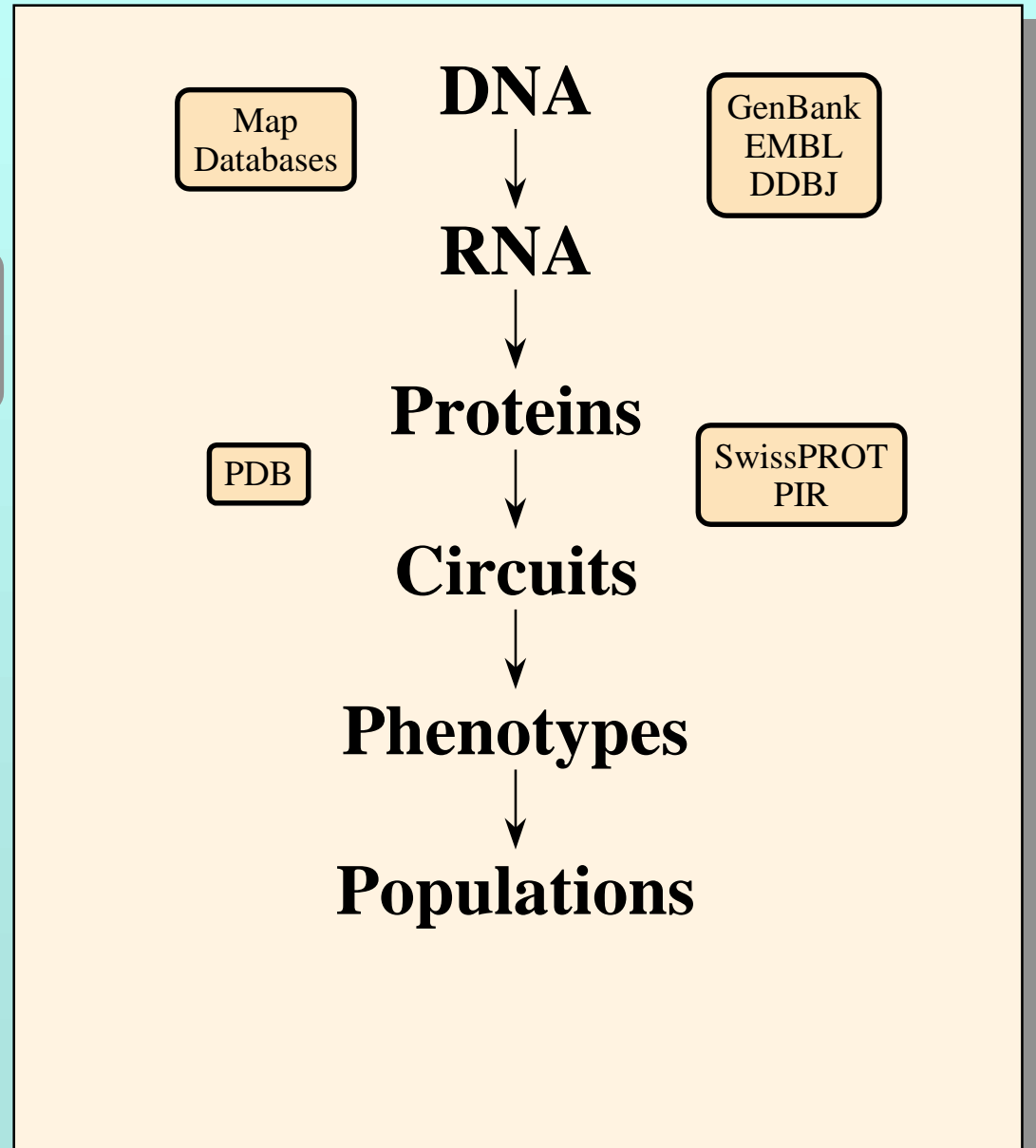
The fundamental dogma of molecular biology is that genes act to create phenotypes through a flow of information from DNA to RNA to proteins, to interactions among proteins (regulatory circuits and metabolic pathways), and ultimately to phenotypes.

Collections of individual phenotypes, of course, constitute a population.



# Fundamental Dogma

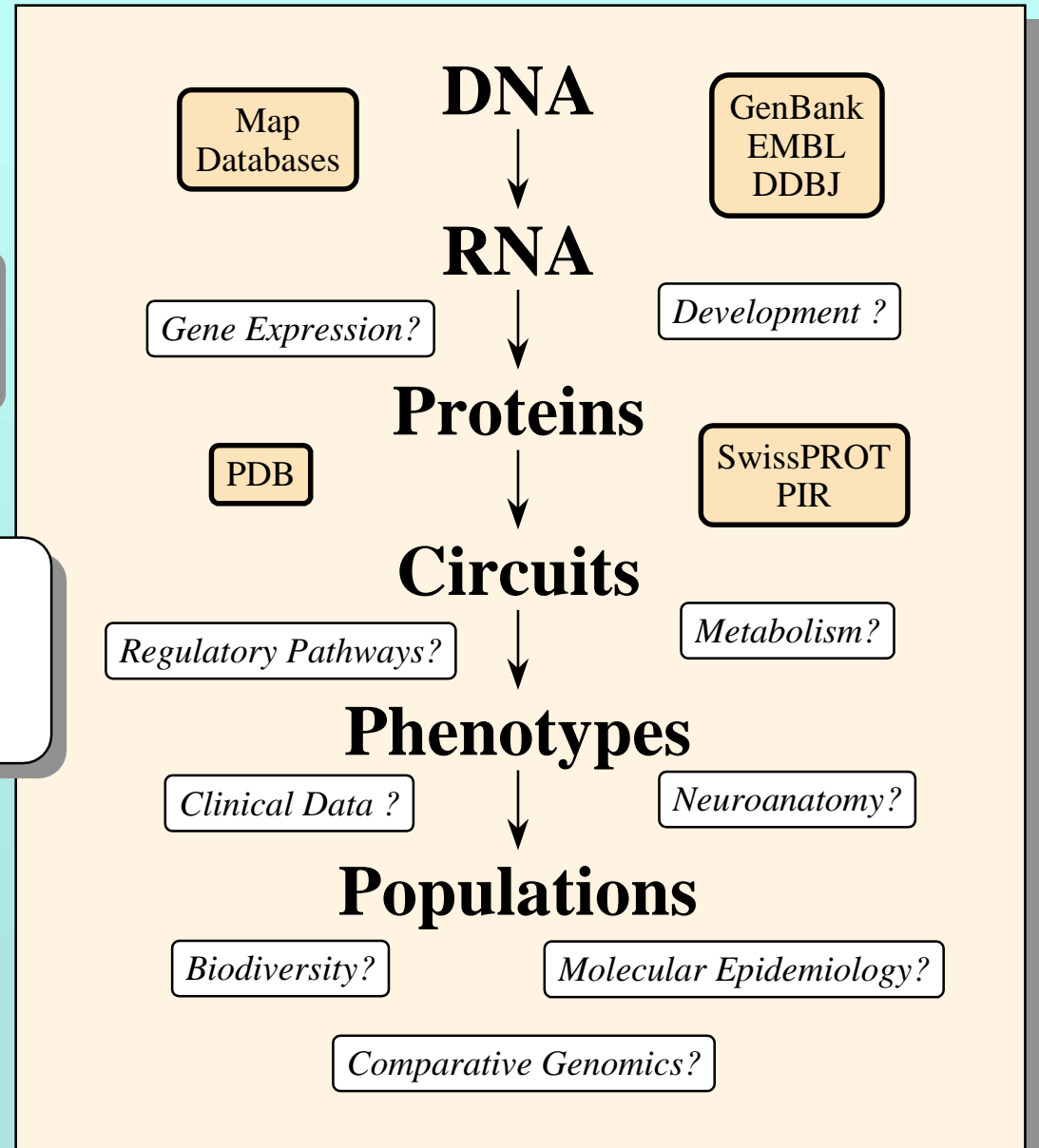
Although a few databases already exist to distribute molecular information,



# Fundamental Dogma

Although a few databases already exist to distribute molecular information,

the post-genomic era will need many more to collect, manage, and publish the coming flood of new findings.

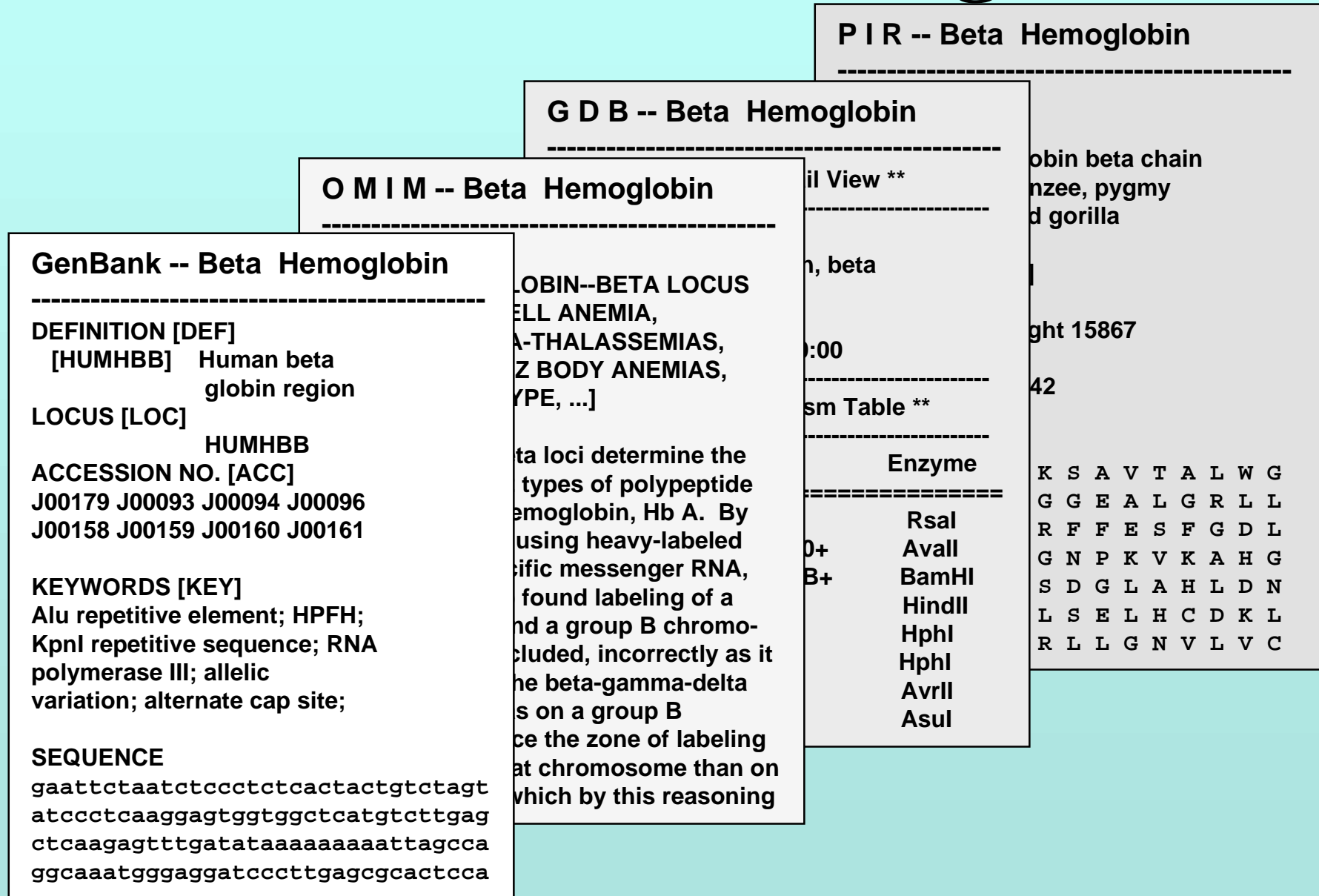


# 21st Century Biology

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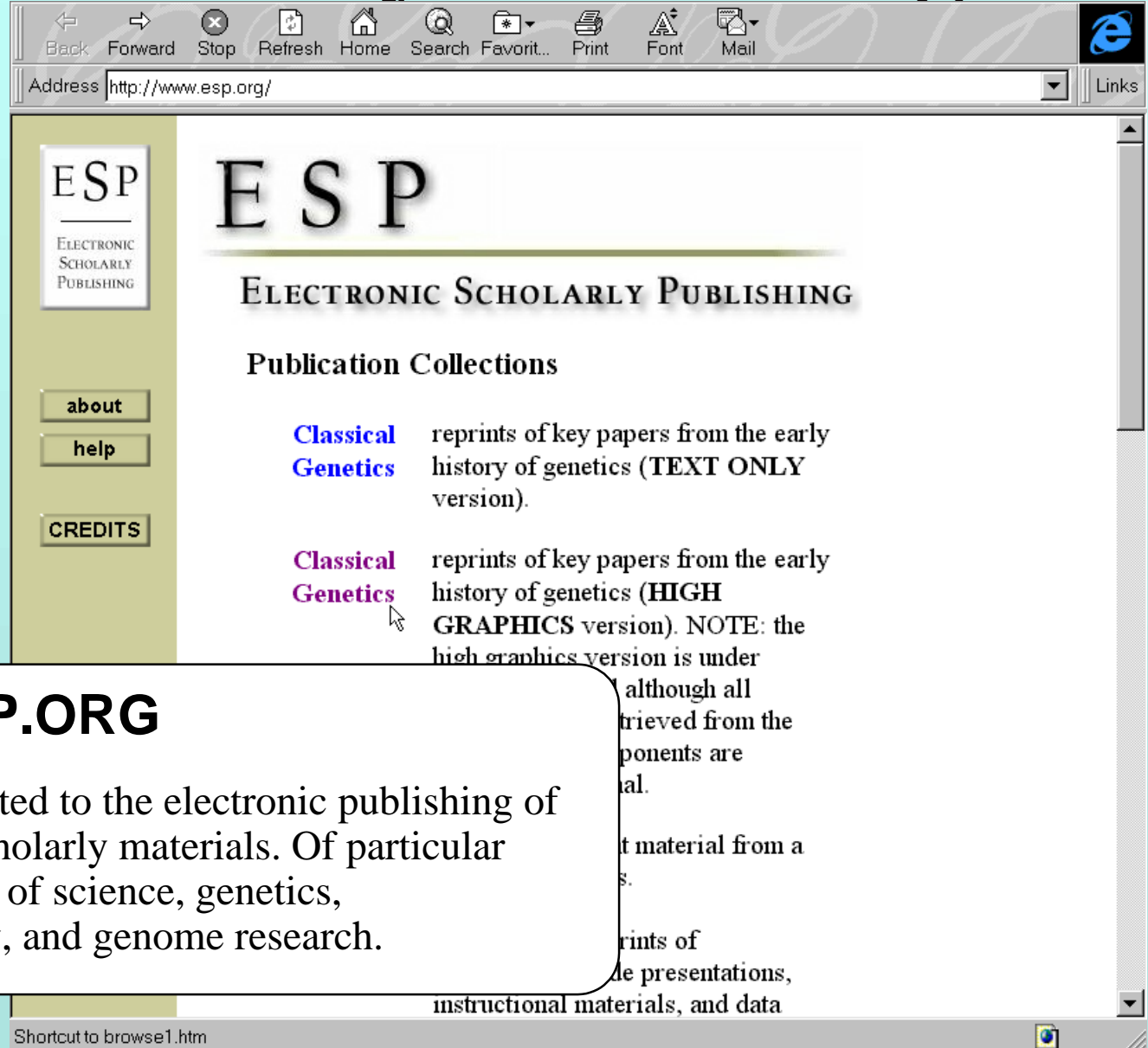
*The Literature*

# Electronic Data Publishing





# Electronic Scholarly Publishing



**HTTP://WWW.ESP.ORG**

The ESP site is dedicated to the electronic publishing of scientific and other scholarly materials. Of particular interest are the history of science, genetics, computational biology, and genome research.

# Electronic Scholarly Publishing

The *Classical Genetics: Foundations* series provides ready access to typeset-quality, electronic editions of important publications that can otherwise be very difficult to find.

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Classical Genetics: Foundations

Early Mendelism

(395,699 bytes; 41 pages, no figures)

Mendel, Gregor. 1865. Experiments in plant hybridization. *Verhandlungen des naturforschenden Vereines in Brünn, Bd. IV für das Jahr 1865, Abhandlungen*, 3-47.

In February and March of 1865, the Brünn Natural History Society in Brünn, Czechoslovakia, heard Gregor Mendel present the results of his investigations into the mechanisms governing inheritance in pea plants. The next year, the work was published as Mendel, Gregor. 1866. "Versuche über Pflanzen Hybriden." *Verhandlungen des naturforschenden Vereines in Brünn*, 4:3-47.

In this remarkable work, Mendel established the foundation for what later became the science of genetics. However, his work was largely ignored when it appeared and Mendel moved on to other things. He died in 1884, never knowing that his experiments in plant hybridization were of any consequence.

His work was rediscovered at the turn of the century and its significance immediately recognized. Genetics, as a formal scientific discipline, exploded into activity in 1900.

Gregor Mendel, 1862

# Electronic Scholarly Publishing

“Hardy” (of Hardy-Weinberg) is a name well known to most students of biology.

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Classical Genetics: Foundations *Early Mendelism*

(19,473 bytes; 1 page, no figures)  [▶ Hardy, G. H. 1908. Mendelian Proportions in a Mixed Population. \*Science, NS. XXVIII: 49-50\*](#)

Every geneticist has heard of the Hardy-Weinberg Law and of Hardy-Weinberg Equilibrium, and nearly all basic biology texts teach that G. H. Hardy played a seminal role in founding population genetics. But, what most biologists don't realize is that Hardy's **total** contribution to biology consisted of a **single** letter to the editor in *Science*. The letter began,

*I am reluctant to intrude in a discussion concerning matters of which I have no expert knowledge, and I should have expected the very simple point which I wish to make to have been familiar to biologists. However, some remarks of Mr. Udry Yule, to which Mr. R. C. Punnett has called my attention, suggest that it may still be worth making*

With that, Hardy offered his "simple point" and then washed his hands of biology. His autobiography, *A Mathematician's Apology*, makes no mention of population genetics.

# Electronic Scholarly Publishing

But how many have read, or even seen, **all** of Hardy's biological writings?

This is it: A single, one-page letter to the editor of *Science*.

The screenshot shows a web browser window with the address bar containing <http://www.esp.org/history/hardy.pdf>. The browser's toolbar includes buttons for Back, Forward, Stop, Refresh, Home, Search, Favorites, Print, Font, and Mail. The document content is a page from *Science*, dated July 10, 1908, volume XXVIII, pages 49-50. The title of the article is "DISCUSSION AND CORRESPONDENCE" and the specific topic is "Mendelian Proportions in a Mixed Population". The text is a letter to the editor by R. C. Punnett, discussing Mendelian inheritance and the concept of a "stable" distribution. The letter is addressed to the Editor of *Science* and mentions a criticism by Mr. Udney Yule. The text discusses the inheritance of brachydactyly and the mathematical models used to describe the distribution of genotypes in a population over generations. The letter concludes by stating that the simple issue of random mating is sufficient to explain the observed results.

JULY 10, 1908

SCIENCE

N. S. Vol. XXVIII:49-50

DISCUSSION AND CORRESPONDENCE

Mendelian Proportions in a Mixed Population

To The Editor of Science: I am reluctant to intrude in a discussion concerning matters of which I have no expert knowledge, and I should have expected the very simple point which I wish to make to have been familiar to biologists. However, some remarks of Mr. Udney Yule, to which Mr. R. C. Punnett has called my attention, suggest that it may still be worth making.

In the *Proceedings of the Royal Society of Medicine* (Vol I., p. 165) Mr. Yule is reported to have suggested, as a criticism of the Mendelian position, that if brachydactyly is dominant "in the course of time one would expect, in the absence of counteracting factors, to get three brachydactylous persons to one normal."

It is not difficult to prove, however, that such an expectation would be quite groundless. Suppose that *Aa* is a pair of Mendelian characters, *A* being dominant, and that in any given generation the numbers of pure dominants (*AA*), heterozygotes (*Aa*), and pure recessives (*aa*) are as  $p:2q:r$ . Finally, suppose that the numbers are fairly large, so that the mating may be regarded as random, that the sexes are evenly distributed among the three varieties, and that all are equally fertile. A little mathematics of the multiplication-table type is enough to show that in the next generation the numbers will be as

this proportion would afterwards have no tendency to decrease.

In a word, there is not the slightest foundation for the idea that a dominant character should show a tendency to spread over a whole population, or that a recessive should tend to die out.

I ought perhaps to add a few words on the effect of the small deviations from the theoretical proportions which will, of course, occur in every generation. Such a distribution as  $p_1:2q_1:r_1$ , which satisfies the condition  $q = p_1r_1$ , we may call a *stable* distribution. In actual fact we shall obtain in the second generation not  $p_1:2q_1:r_1$  but a slightly different distribution  $p_2:2q_2:r_2$ , which is not "stable." This should, according to theory, give us in the third generation a "stable" distribution  $p_3:2q_3:r_3$ , also differing from  $p_1:2q_1:r_1$ ; and so on. The sense in which the distribution  $p_1:2q_1:r_1$  is "stable" is this, that if we allow for the effects of casual deviations in any subsequent generation, we should, according to theory, obtain at the next generation a new "stable" distribution differing but slightly from the original distribution.

I have, of course, considered only the very simplest hypotheses possible. Hypotheses other than [sic] that of purely random mating will give different results, and, of course, if, as appears to be the case sometimes, the character is not independent of that of sex, or has an influence on fertility, the whole question may be greatly complicated. But such complications seem to be irrelevant to the simple issue

Page 1 of 1 | 128% | 8.5 x 11 in

# Electronic Scholarly Publishing

The screenshot shows a web browser window with the address bar containing <http://www.esp.org/books/darwin/beagle/>. The browser's toolbar includes buttons for Back, Forward, Stop, Refresh, Home, Search, Favorites, Print, Font, and Mail. The website content is displayed in a black frame. On the left side of the frame, there is a logo for ESP (Electronic Scholarly Publishing) and a vertical menu with buttons for 'about', 'help', 'search', and 'CREDITS'. The main content area of the page is white and features the following text: 'Charles Darwin' at the top left, 'Voyage of the Beagle' at the top right, 'Electronic Scholarly Publishing' in purple italics, 'HTML Edition' in black italics, 'THE VOYAGE OF THE BEAGLE' in bold black uppercase, '2nd Edition' in black italics, 'BY CHARLES DARWIN' in bold black uppercase, 'From The Harvard Classics Volume 29', 'Copyright, 1909', 'P. F. Collier & Son, New York', and 'Table of Contents' in purple italics at the bottom. A mouse cursor is positioned over the 'Table of Contents' link.

<http://www.esp.org/books/darwin/beagle>

Entire monographs can be made instantly available to readers world-wide..

# Electronic Scholarly Publishing

Today's computer technology was nearly unimaginable just ten years ago. The technology of ten years from now will also bring many surprises.

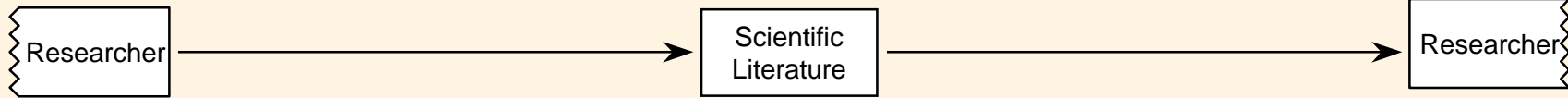
How is it that IT can maintain such an amazing rate of sustained change?

And what, if any, are the implications of that rate of change for biology?

The screenshot shows a web browser window with the address bar containing <http://www.esp.org/books/darwin/beagle/>. The browser's toolbar includes buttons for Back, Forward, Stop, Refresh, Home, Search, Favorites, Print, Font, and Mail. The main content area displays the title page for 'THE VOYAGE OF THE BEAGLE' by Charles Darwin, with the subtitle 'Contents'. The text is centered and includes a list of chapters from I to XIV, such as 'ST. JAGO -- CAPE DE VERD ISLANDS' and 'RIO DE JANEIRO'. A mouse cursor is visible over the text 'RIO DE JANEIRO'. On the left side of the browser window, there is a vertical navigation menu with buttons for 'about', 'help', 'search', and 'CREDITS'. The ESP logo is visible in the top left corner of the page content. At the bottom left of the page, there is a copyright notice: 'Copyright, 1996, 1997, E.S.P.'. The browser's status bar at the bottom shows 'Done'.

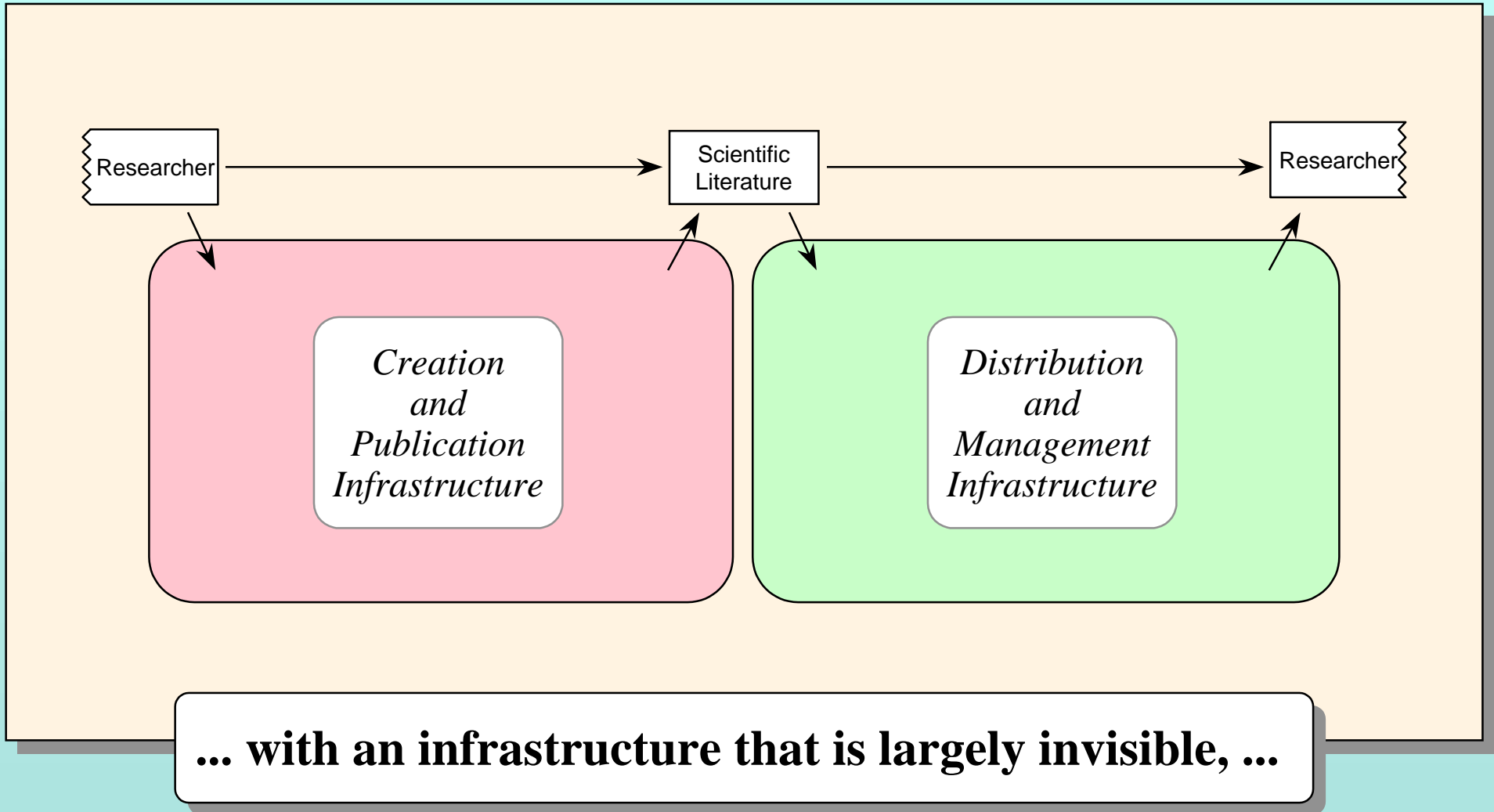
# Traditional Publishing

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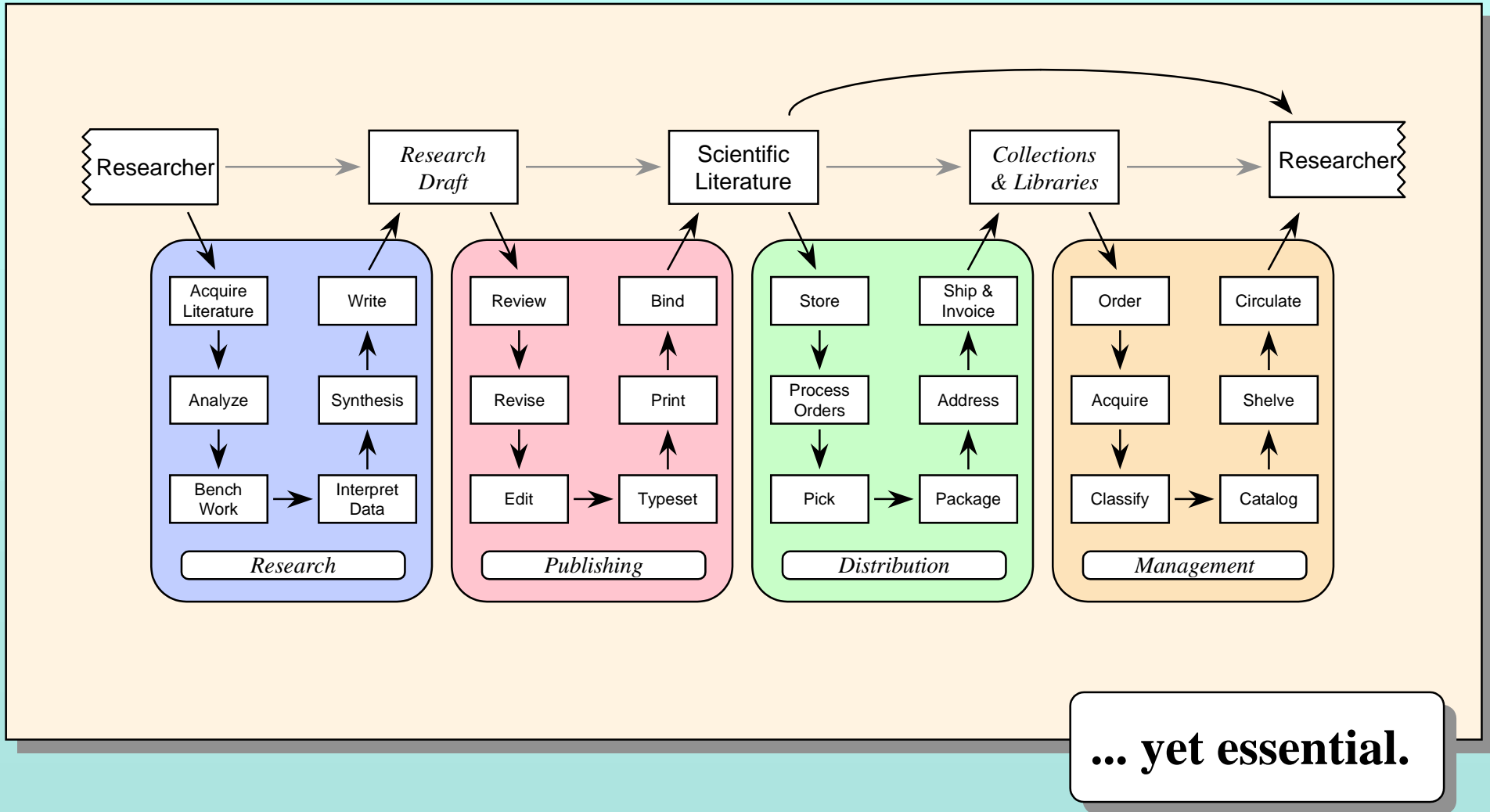
**Print publication seems straightforward, ...**

# Traditional Publishing

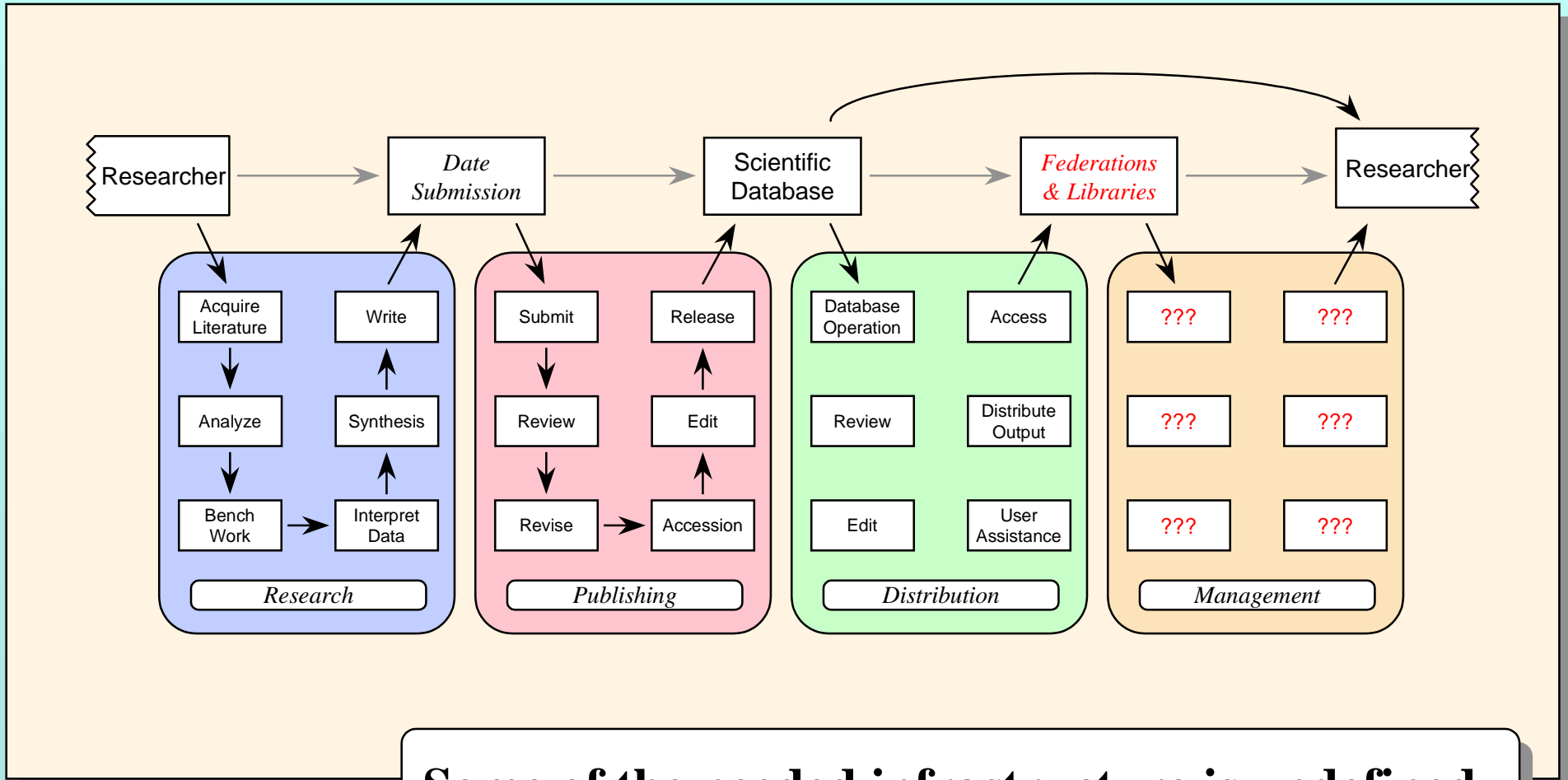




# Traditional Publishing



# Electronic Publishing



# 21st Century Biology

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*The People*

# Human Resources Issues

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- Reduction in need for non-IT staff

# Human Resources Issues

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- Reduction in need for non-IT staff
- Increase in need for IT staff, especially “information engineers”

# Human Resources Issues

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- Reduction in need for non-IT staff
- Increase in need for IT staff, especially “information engineers”

In modern biology, a general trend is to convert expert work into staff work and finally into computation. New expertise is required to design, carry out, and interpret continuing work.

# Human Resources Issues

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**Elbert Branscomb:** “You must recognize that some day you may need as many computer scientists as biologists in your labs.”

# Human Resources Issues

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**Elbert Branscomb:** “You must recognize that some day you may need as many computer scientists as biologists in your labs.”

**Craig Venter:** “At TIGR, we already have twice as many computer scientists on our staff.”

Exchange at DOE workshop on high-throughput sequencing.



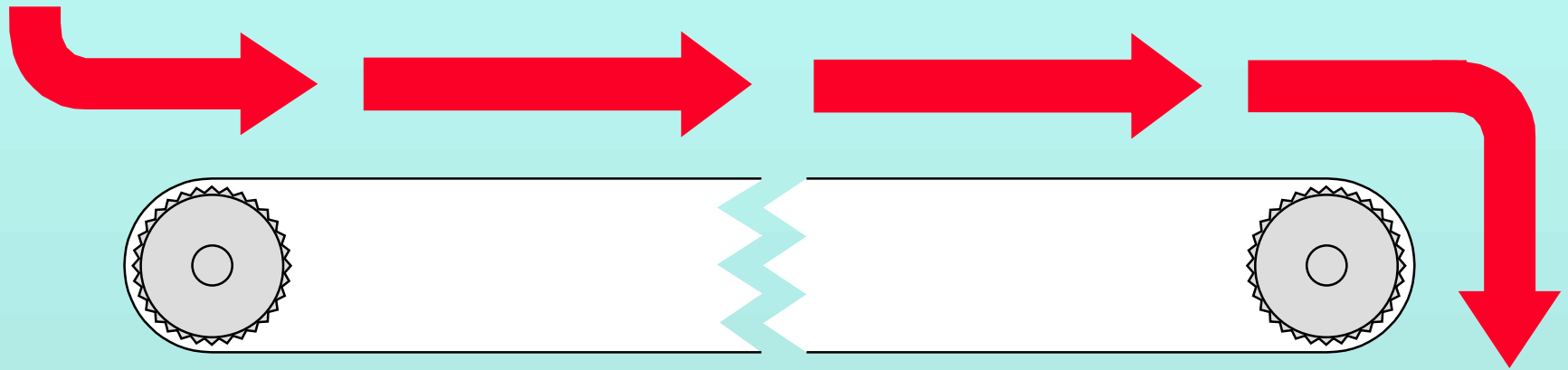
# New Discipline of Informatics

# What is Informatics?

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Computer  
Science  
Research

----- Informatics -----



Biological  
Application  
Programs

# What is Informatics?

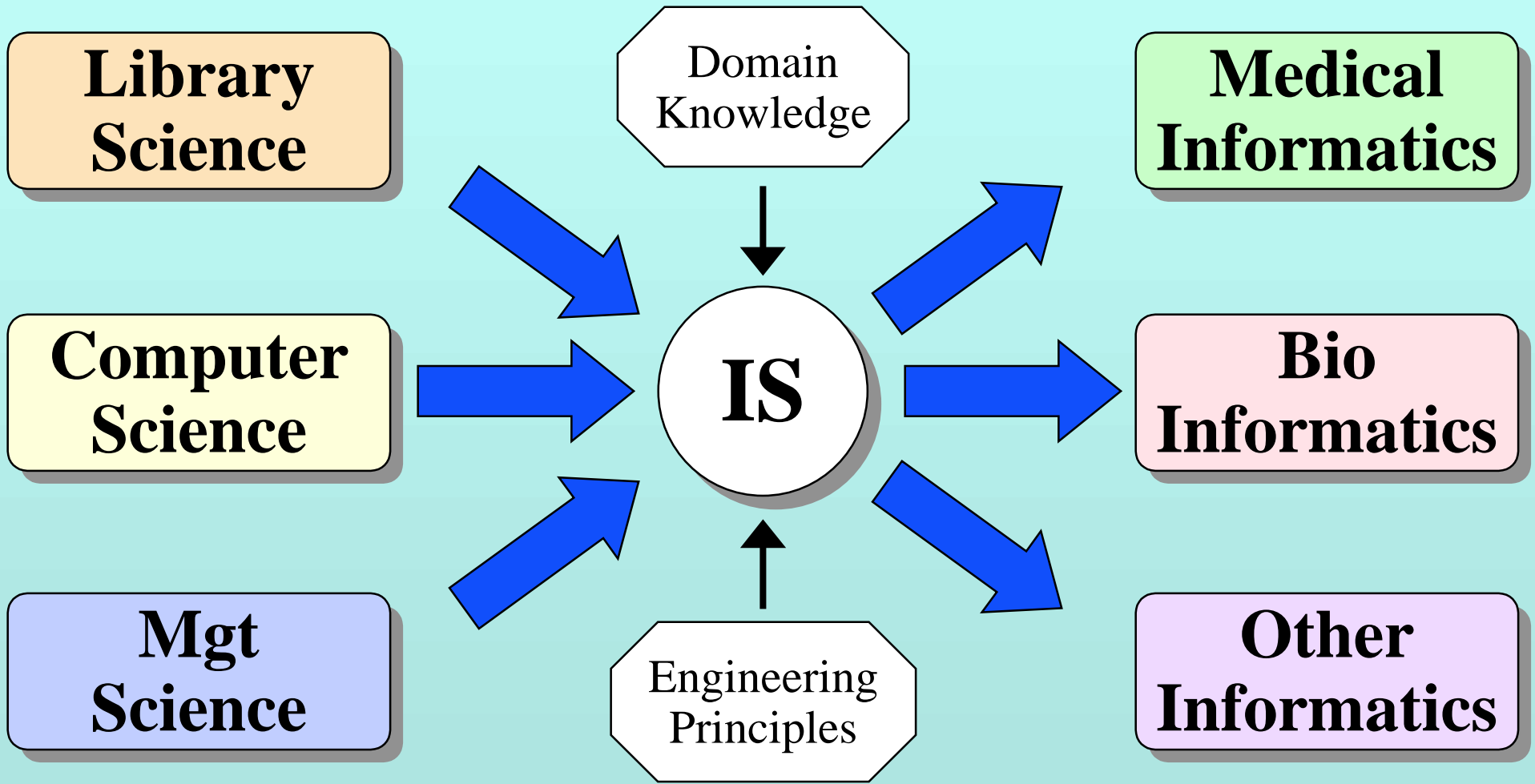
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Informatics combines expertise from:

- *domain science (e.g., biology)*
- *computer science*
- *library science*
- *management science*

All tempered with an engineering mindset...

# What is Informatics?



# Engineering Mindset

---

Engineering is often defined as the use of scientific knowledge and principles for practical purposes. While the original usage restricted the word to the building of roads, bridges, and objects of military use, today's usage is more general and includes chemical, electronic, and even mathematical engineering.

Parnas, David Lorge. 1990. *Computer*, 23(1):17-22.

# Engineering Mindset

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... or even information engineering.

# Engineering Mindset

---

Engineering education ... stresses finding good, as contrasted with workable, designs. Where a scientist may be happy with a device that validates his theory, an engineer is taught to make sure that the device is efficient, reliable, safe, easy to use, and robust.

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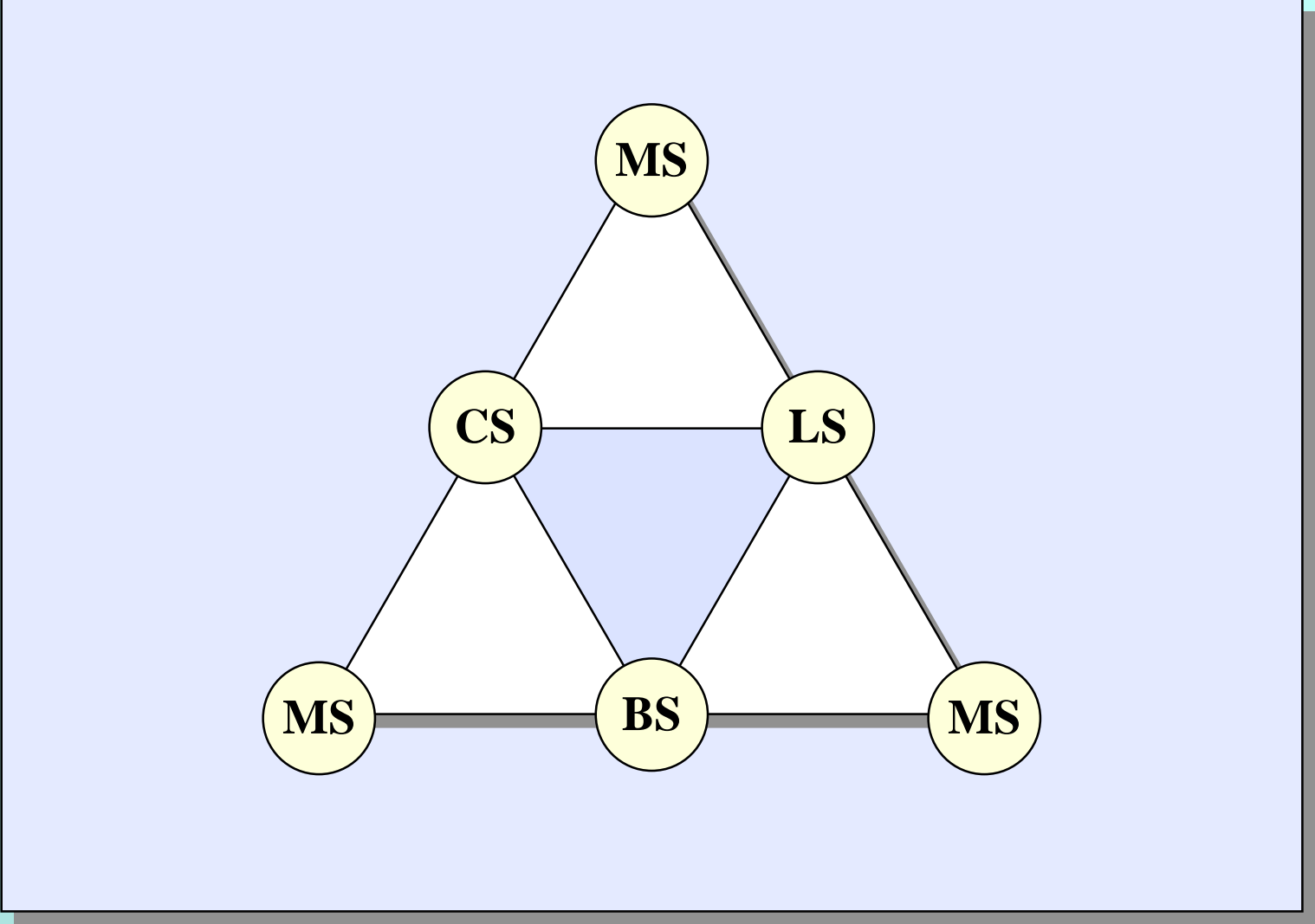
Parnas, David Lorge. 1990. *Computer*, 23(1):17-22.

The assembly of working, robust systems, on time and on budget, is the key requirement for a federated information infrastructure for biology.



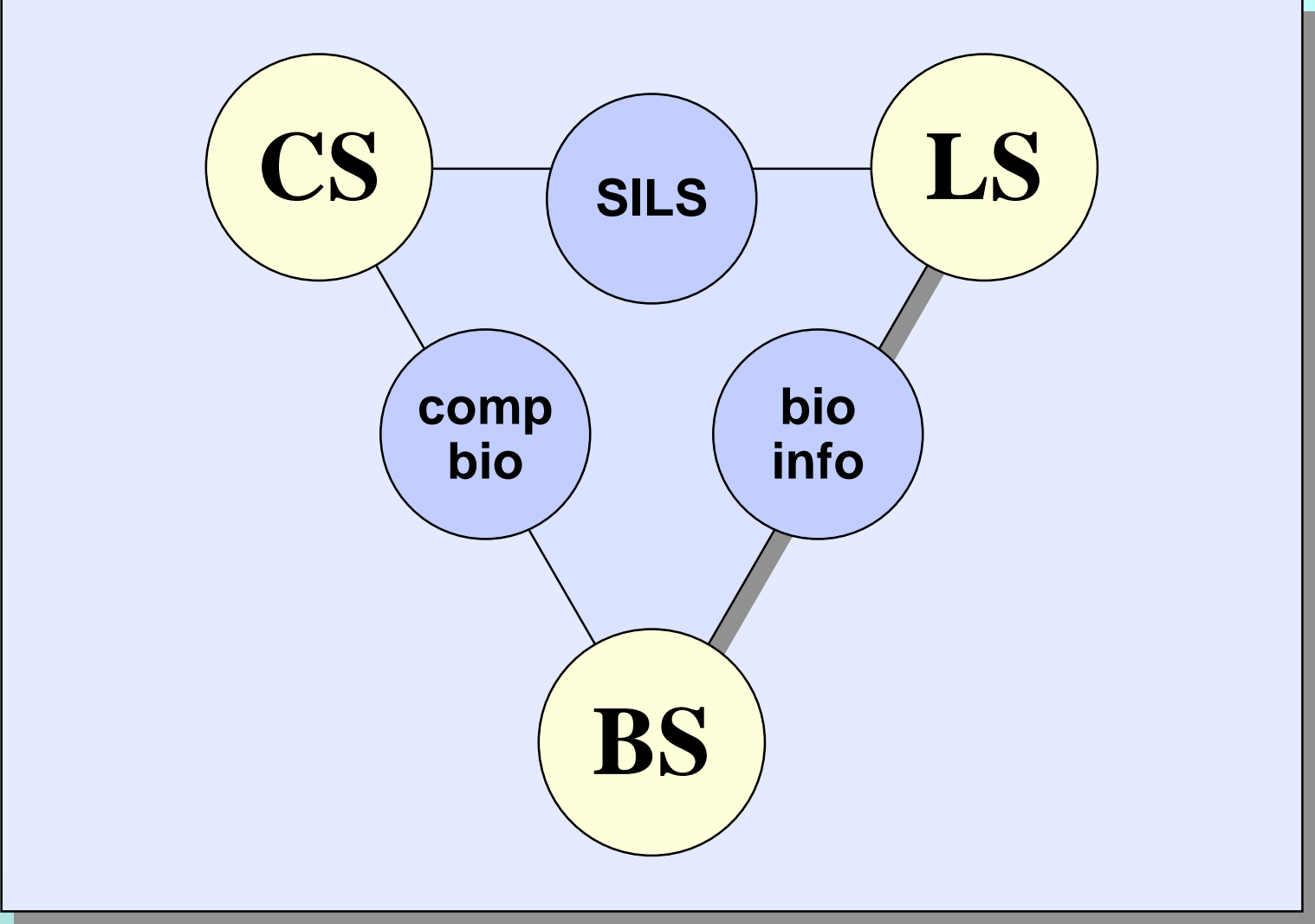
# Informatics Triangle

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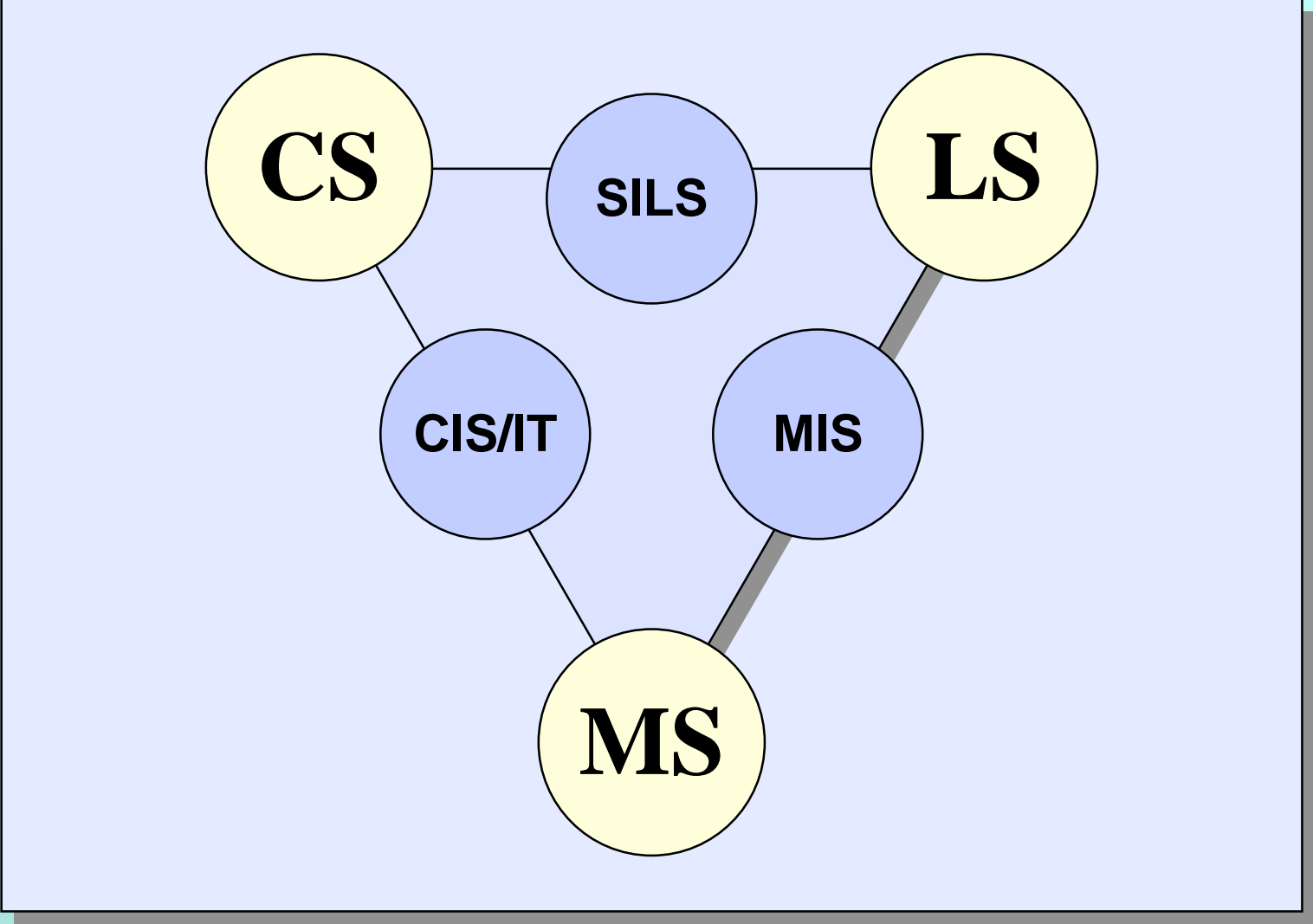
# Informatics Triangle

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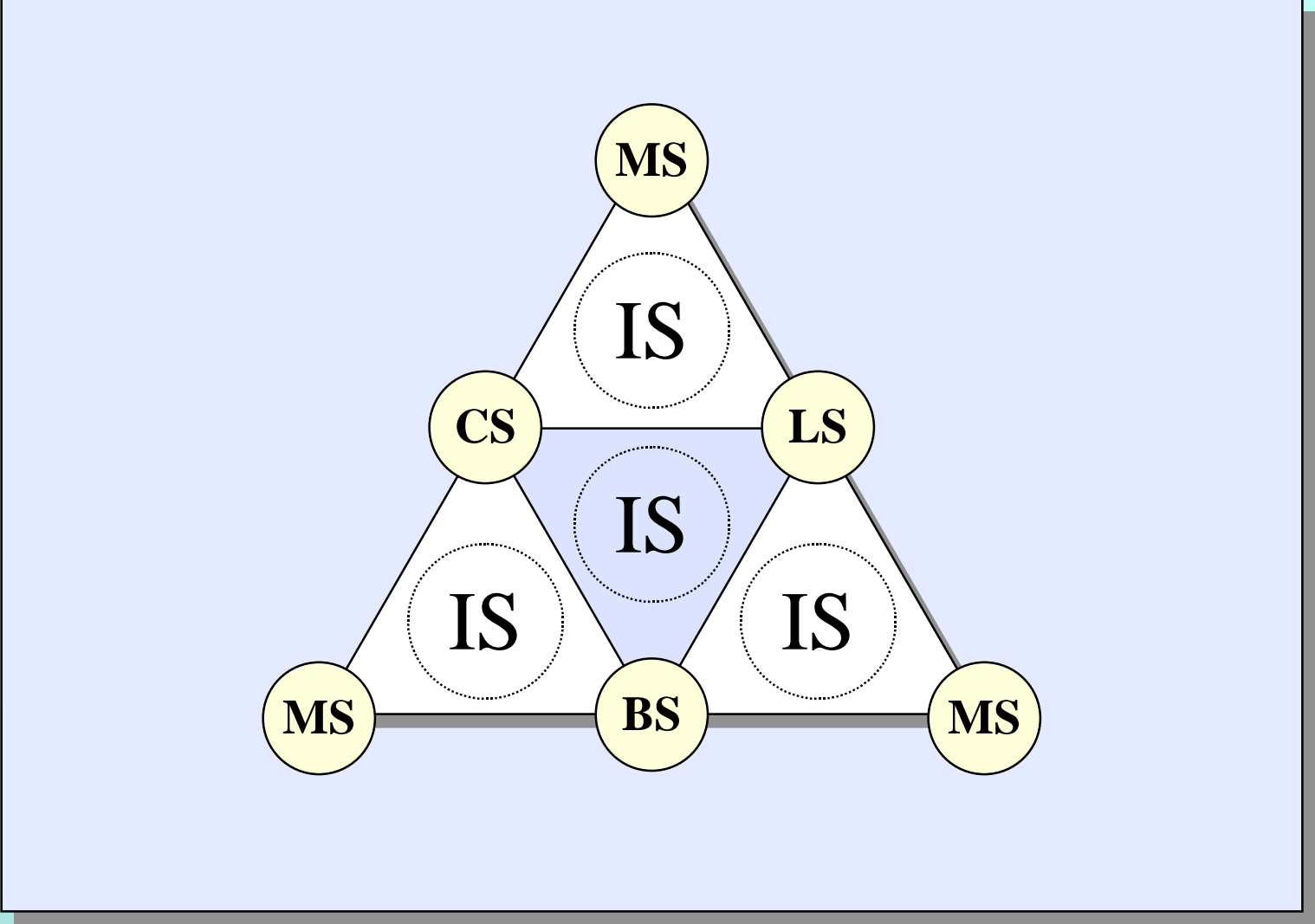
# Informatics Triangle

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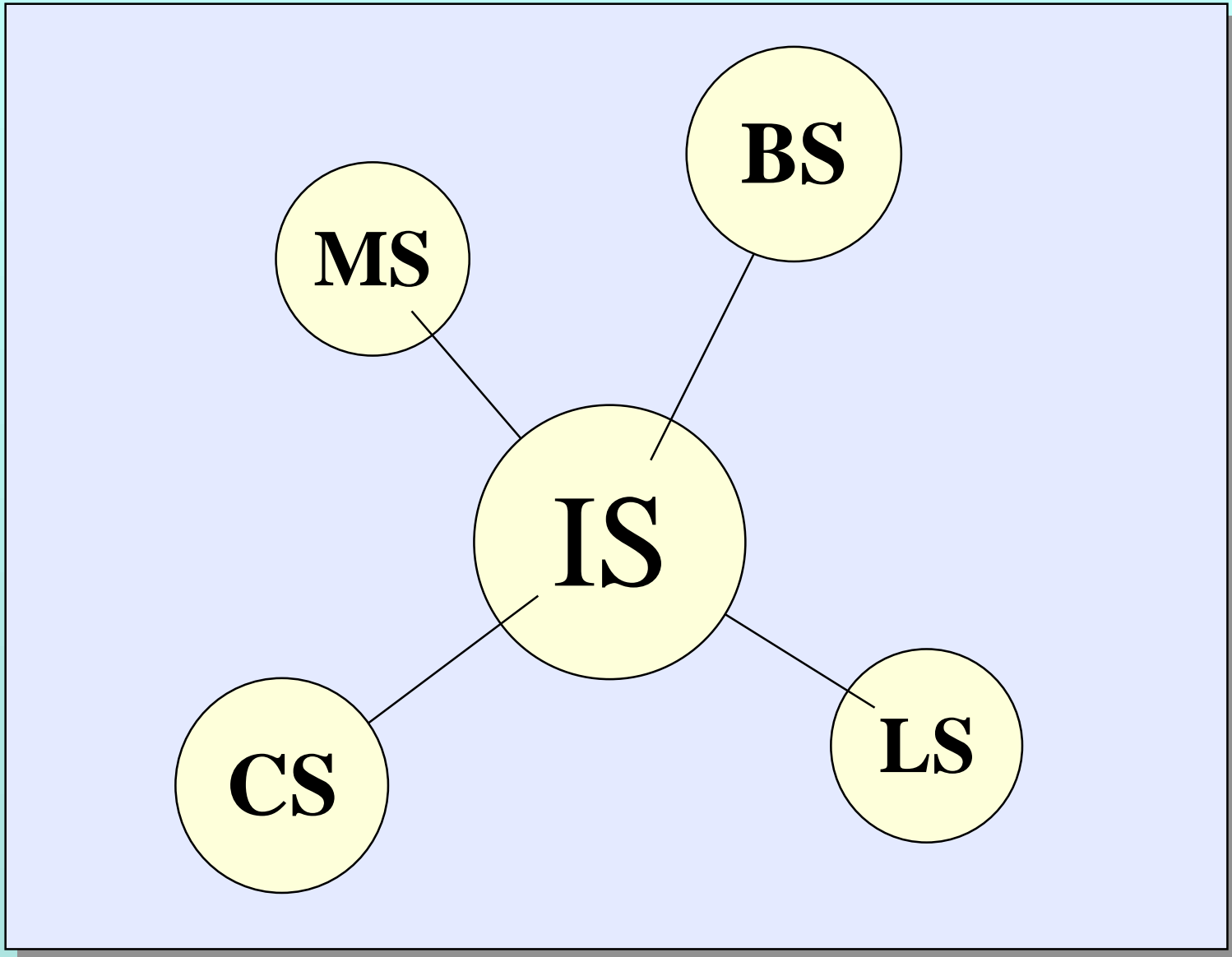
# Informatics Triangle

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# What is Informatics?

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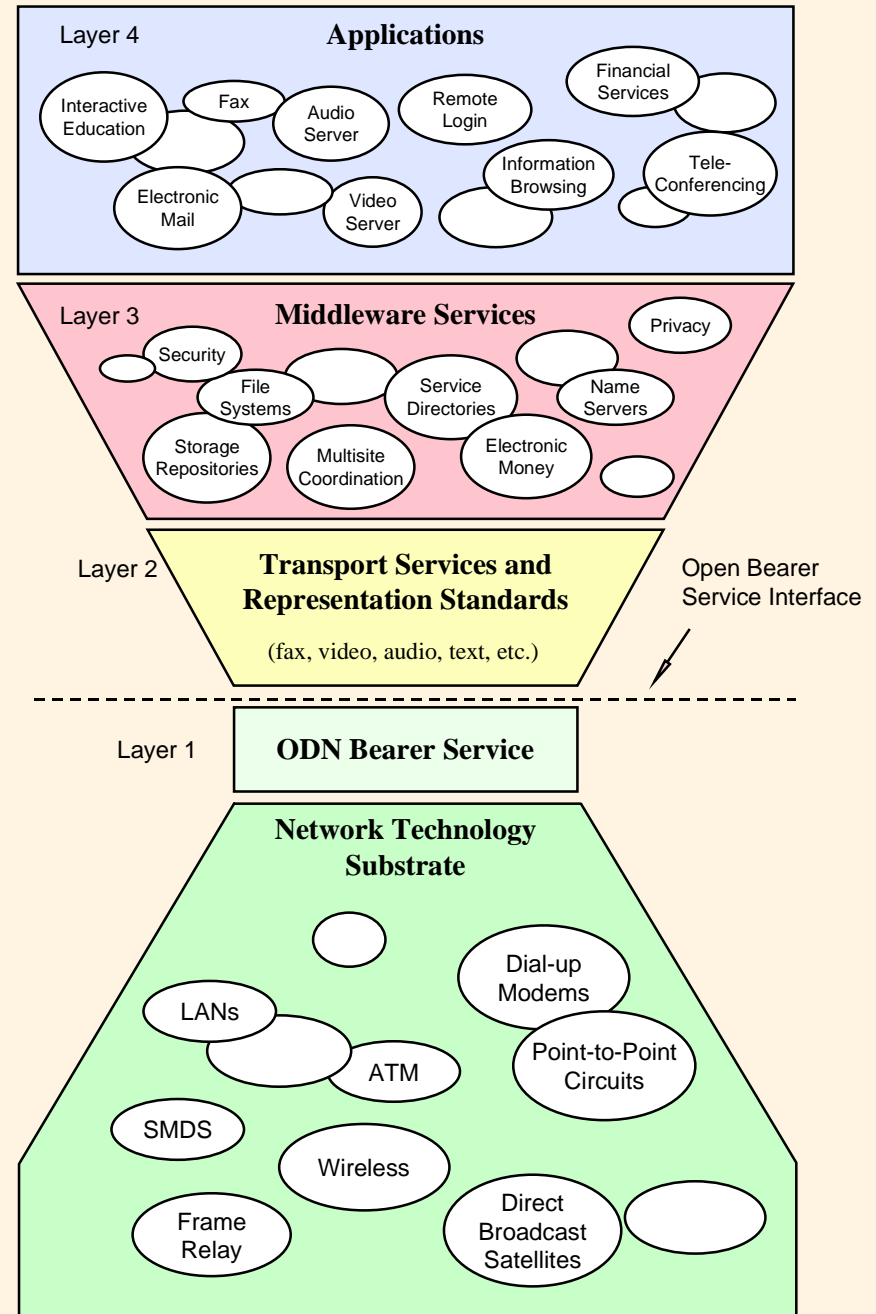
# Federated Information Infrastructure

# National Information Infrastructure

|         | commercial uses |       | non-commercial uses |     |  |
|---------|-----------------|-------|---------------------|-----|--|
|         | ETC             | other | Edu                 | Lib | Res  |
| analog  |                 |       |                     |     |  |
| digital |                 |       |                     |     |  |

# ODN Model

A recent NRC report, *Realizing the Information Future*, laid out a vision of an Open Data Network model, in which any information appliance could be operated over generic networking protocols...

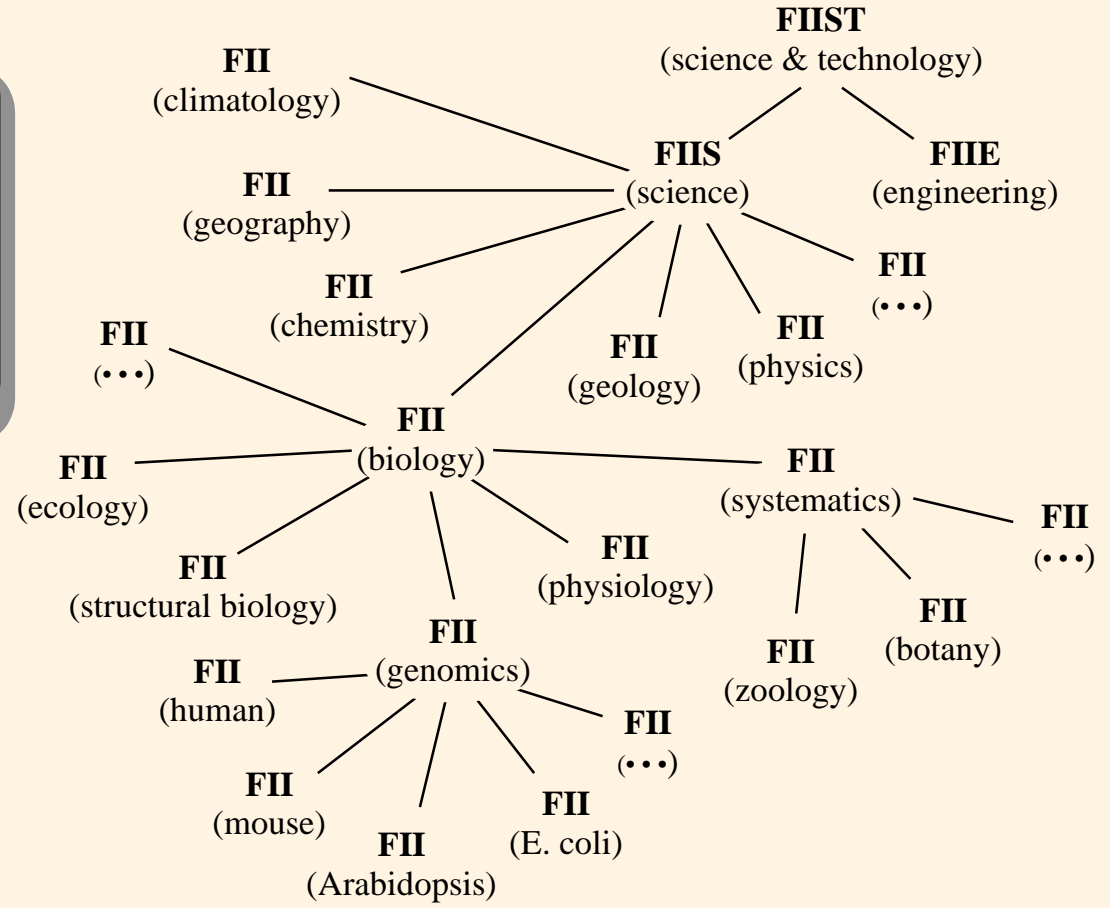




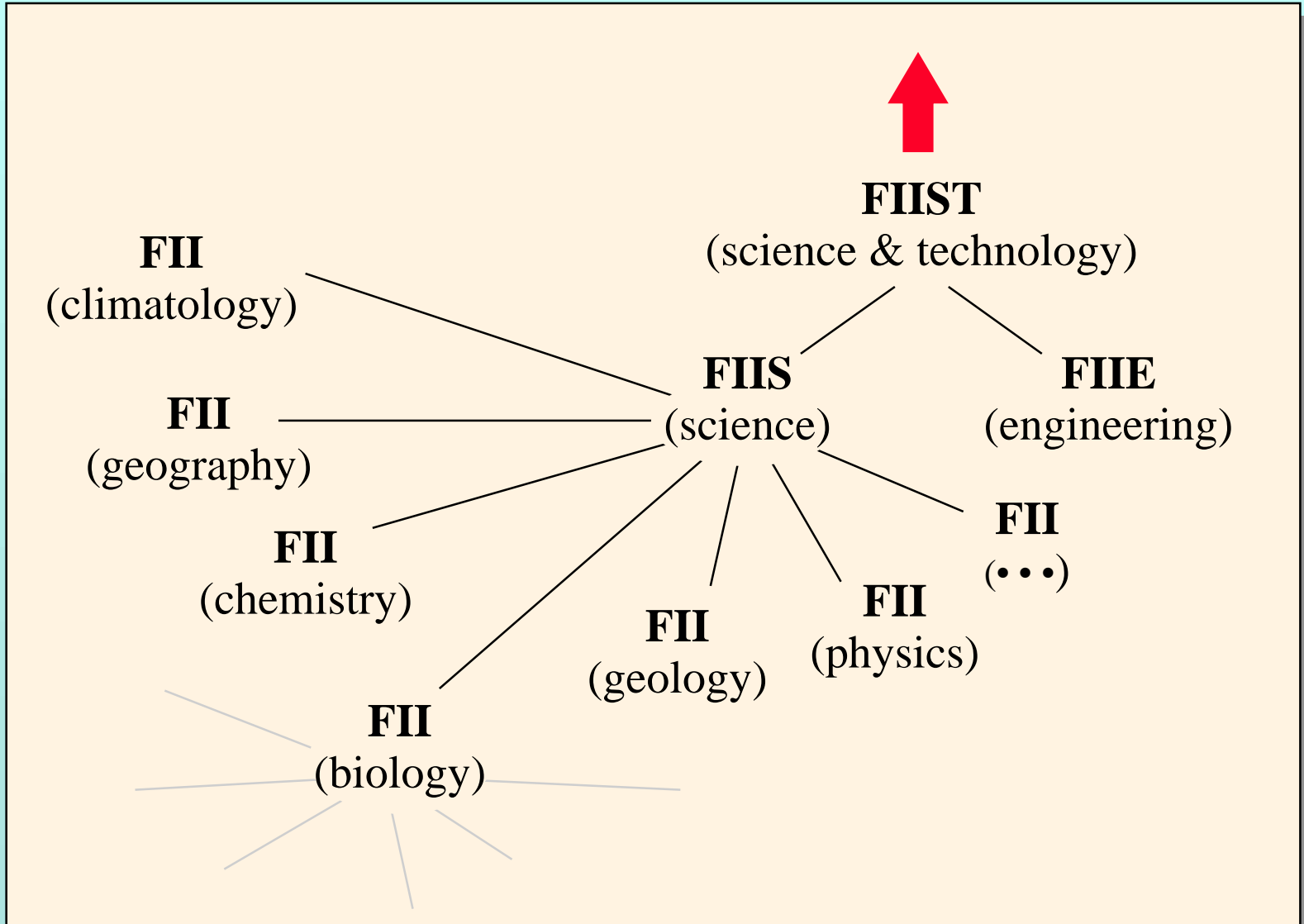
# FIIST & NII

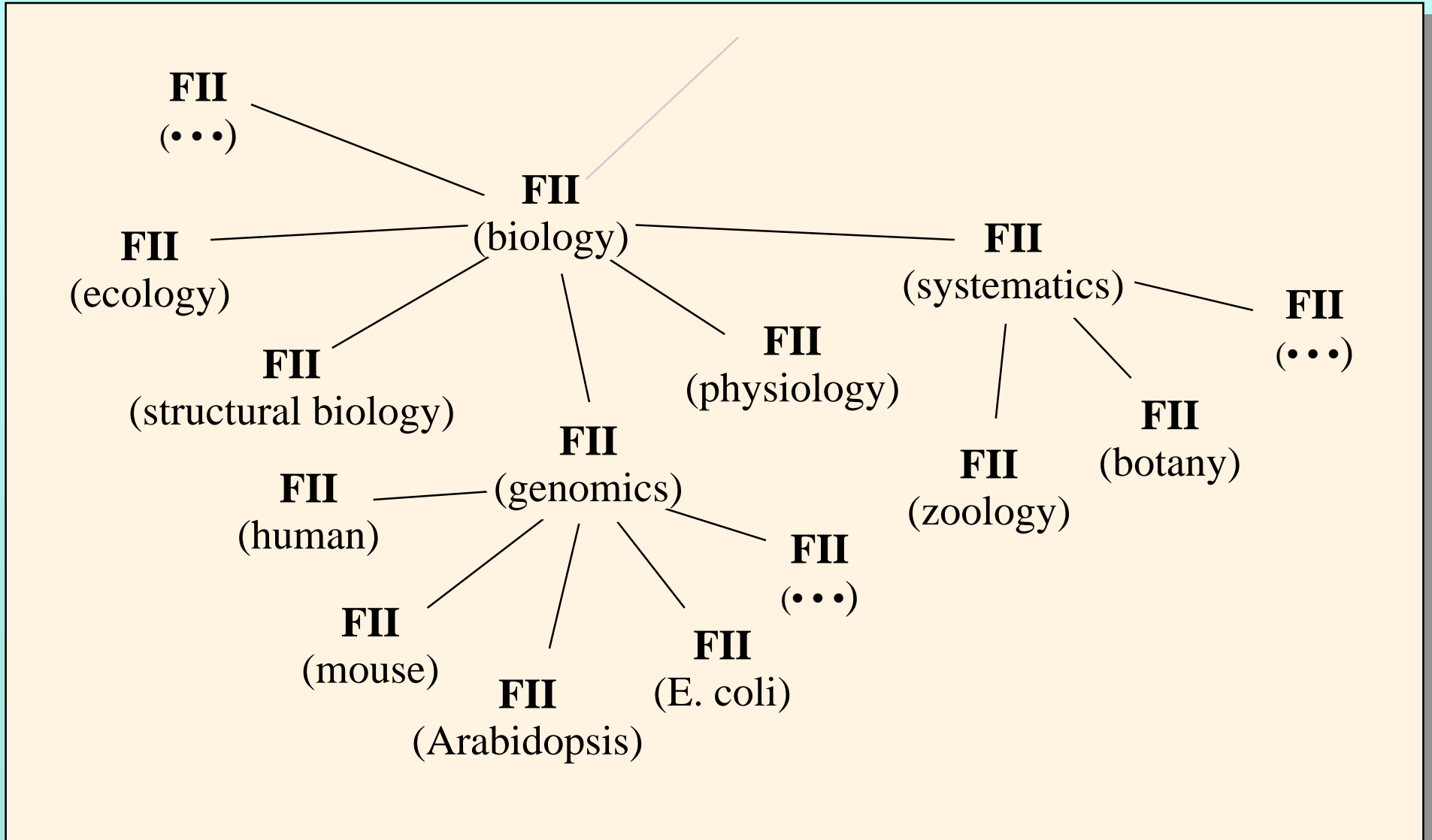
The research component of the NII contains a Federated Information Infrastructure for Science and Technology..

|         | commercial uses |       | non-commercial uses |     |      |
|---------|-----------------|-------|---------------------|-----|------|
|         | ETC             | other | Edu                 | Lib | Res  |
| analog  |                 |       |                     |     |      |
| digital |                 |       |                     |     | ◆◆◆◆ |



# FIIST





# Public Funding of Databases

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## Stand-alone Criteria:

- Is there a need?
- Will this meet the need?
- Can they do it?
- Is it worth it?

# Public Funding of Databases

---

## Global Criteria:

- Does it adhere to standards?
- Will it interoperate?
- Is there commitment to federation?
- Is it worth it?

# Information Resources and the GII

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## Guiding Principles:

- Global value explosion
- Componentry
- Anonymous interoperability
- Technical scalability
- Social scalability
- Value additivity

# Funding for Bio-Information Infrastructure

# Call for Change

---

Among the many new tools that are or will be needed (for 21st-century biology), some of those having the highest priority are:

- bioinformatics
- computational biology
- functional imaging tools using biosensors and biomarkers
- transformation and transient expression technologies
- nanotechnologies

*Impact of Emerging Technologies on the Biological Sciences: Report of a Workshop.* NSF-supported workshop, held 26-27 June 1995, Washington, DC.



# The Problem

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- IT moves at “Internet Speed” and responds rapidly to market forces.

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# The Problem

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- IT moves at “Internet Speed” and responds rapidly to market forces.
- IT will play a central role in 21st Century biology.
- Current levels of support for public bio-information infrastructure are too low.
- Reallocation of federal funding is difficult, and subject to political pressures.
- Federal-funding decision processes are ponderously slow and inefficient.

# **Federal Funding of Bio-Databases**

---

**The challenges:**

# Federal Funding of Bio-Databases

## **The challenges:**

- providing adequate funding levels

# Federal Funding of Bio-Databases

## **The challenges:**

- providing adequate funding levels
- making timely, efficient decisions



# IT Budgets

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*A Reality Check*

# Rhetorical Question

---

**Which is likely to be more complex:**

- identifying, documenting, and tracking the whereabouts of **all parcels** in transit in the US at one time
- identifying, documenting, and analyzing the structure and function of **all individual genes in all economically significant organisms**; then analyzing **all significant gene-gene and gene-environment interactions** in those organisms and their environments

# Business Factoids

---

## United Parcel Service:

- uses two redundant 3 Terabyte (yes, 3000 GB) databases to track all packages in transit.
- has 4,000 full-time employees dedicated to IT
- spends one billion dollars per year on IT
- has an income of 1.1 billion dollars, against revenues of 22.4 billion dollars

# Business Comparisons

| Company                | Revenues        | IT Budget     | Pct     |
|------------------------|-----------------|---------------|---------|
| Chase-Manhattan        | 16,431,000,000  | 1,800,000,000 | 10.95 % |
| AMR Corporation        | 17,753,000,000  | 1,368,000,000 | 7.71 %  |
| Nation's Bank          | 17,509,000,000  | 1,130,000,000 | 6.45 %  |
| Sprint                 | 14,235,000,000  | 873,000,000   | 6.13 %  |
| IBM                    | 75,947,000,000  | 4,400,000,000 | 5.79 %  |
| MCI                    | 18,500,000,000  | 1,000,000,000 | 5.41 %  |
| Microsoft              | 11,360,000,000  | 510,000,000   | 4.49 %  |
| United Parcel          | 22,400,000,000  | 1,000,000,000 | 4.46 %  |
| Bristol-Myers Squibb   | 15,065,000,000  | 440,000,000   | 2.92 %  |
| Pfizer                 | 11,306,000,000  | 300,000,000   | 2.65 %  |
| Pacific Gas & Electric | 10,000,000,000  | 250,000,000   | 2.50 %  |
| Wal-Mart               | 104,859,000,000 | 550,000,000   | 0.52 %  |
| K-Mart                 | 31,437,000,000  | 130,000,000   | 0.41 %  |

# Federal Funding of Biomedical-IT

## Appropriate funding level:

- approx. 5-10% of research funding
- *i.e.*, 1 - 2 **billion** dollars per year

### Source of estimate:

- Experience of IT-transformed industries.
- Current support for IT-rich biological research.

Conference on  
**BIOLOGICAL INFORMATICS**

6-8 July 1998

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Australian Academy of Science, Canberra, Australia

# Conference on Biological Informatics

## Conference Sessions

- Overview of Biological Informatics
- Biodiversity Informatics
- Environmental Informatics
- Molecular Informatics
- Medical / Neuroinformatics
- Teaching and Training in Informatics

**Extras**



# Slides:

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**<http://www.esp.org/rjr/canberra.pdf>**

**Extras**

# Basics

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*Business 101*

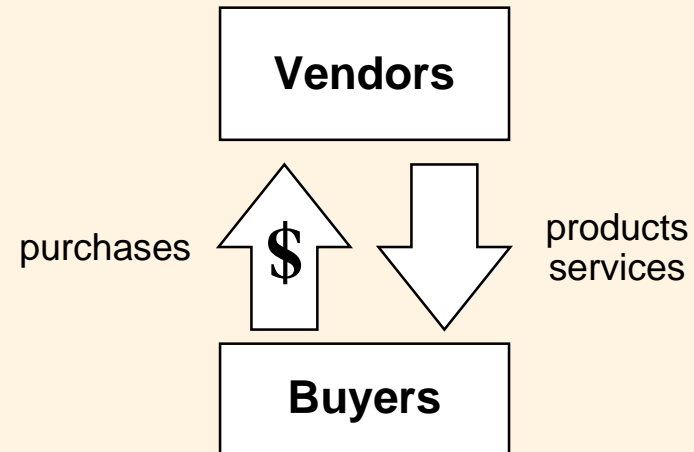
# Market Forces

In a simple market economy, vendors try to anticipate the needs of buyers and offer products and services to meet those needs.

Real users decide whether or not to buy a product or service, depending upon whether or not it meets a real need at a reasonable price.

## Business 101 Insight:

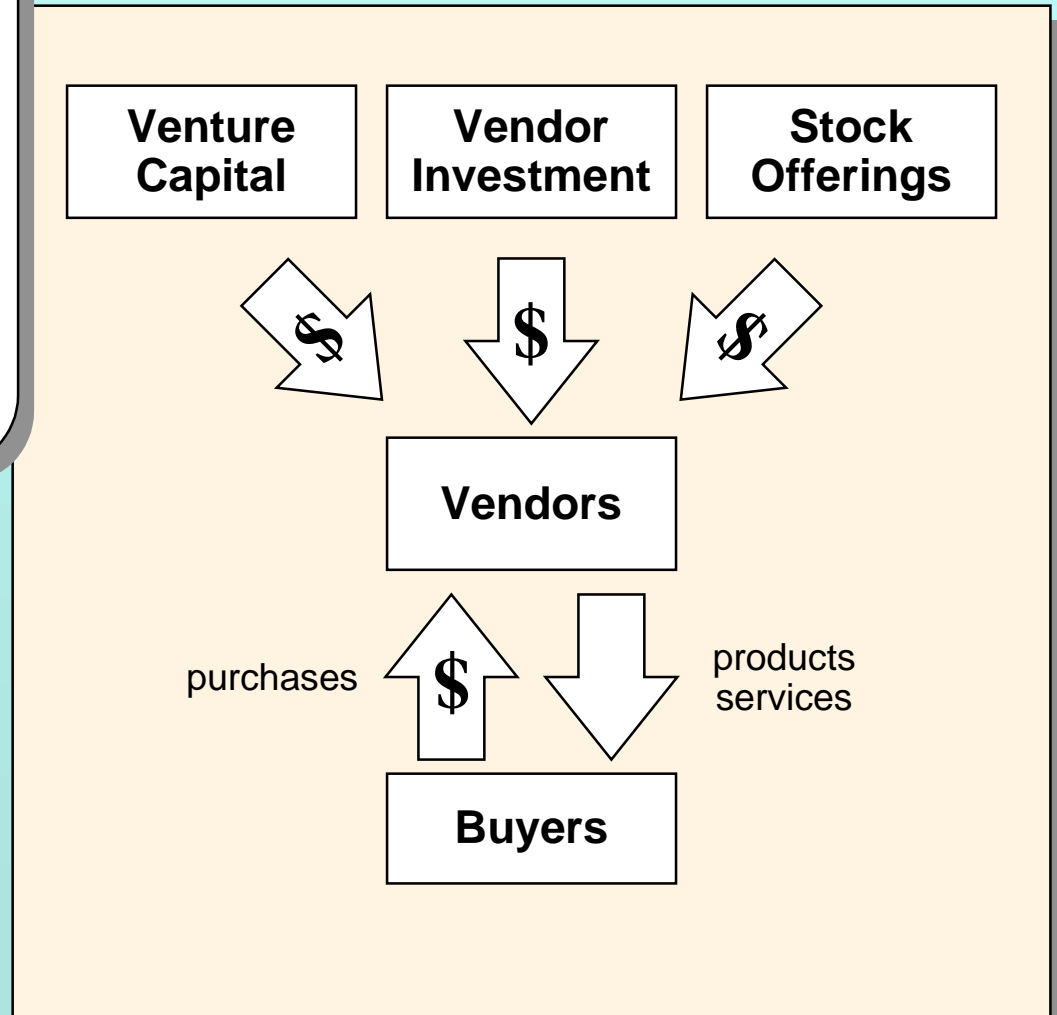
Successful vendors target a niche and excel at meeting the needs of that niche.



# Market Forces

Funding to initiate the development of products and services come from investors, not from buyers.

Investors decide whether or not to provide start-up funding based upon the estimated ability of the vendor to create products and services that will meet real needs at competitive prices.

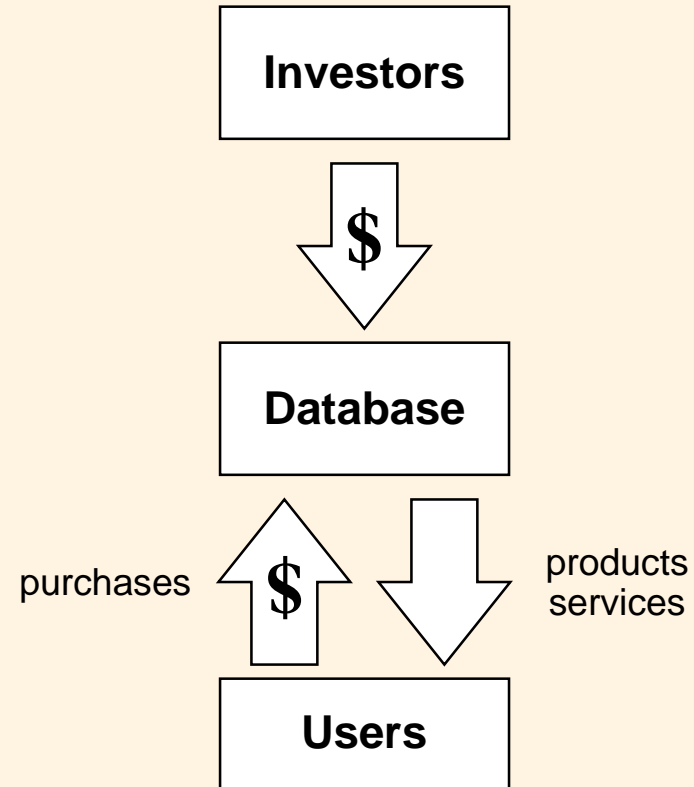


# Federal Funding

If biological databases were driven by market forces, individual users would choose what services they need and individual database providers would choose what services to make available.

Investors would provide start-up money on the likelihood of successful products and services being developed.

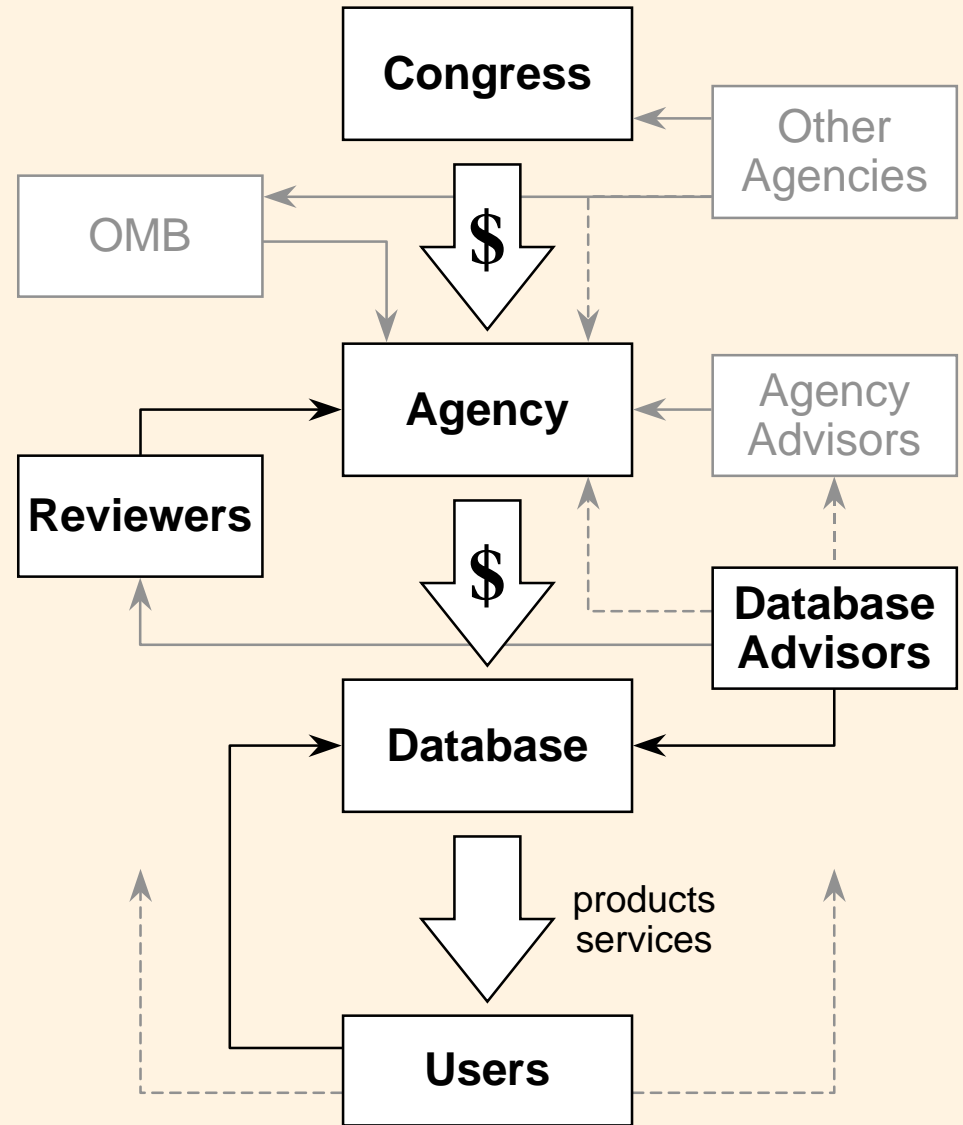
Ultimate success would depend on meeting the needs of real users. Decisions could be made rapidly, in response to changing needs and emerging opportunities.



# Federal Funding

Instead, funding decisions for grant-supported biological databases can follow a ponderously slow course, with almost no opportunity for real-time input from real users.

Even with the best of intentions at all levels, this process is slow, inefficient, risk-averse, and non-responsive to the real and changing needs of users.



# Federal Funding of Bio-Databases

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## Possible solutions:

- increase the direct support of federal service organizations providing information infrastructure (*e.g.*, NCBI).
- reduce support for investigator-initiated, grant-funded public database projects.
- create market forces, initially through subsidization, later simply through direct support for affected science (*e.g.*, NSFnet into internet).



# Federal Funding of Bio-Databases

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## Creating market forces:

- stop supporting the supply side of biodatabases through slow, inefficient processes.

# Federal Funding of Bio-Databases

---

## Creating market forces:

- stop supporting the supply side of biodatabases through slow, inefficient processes.
- start supporting the demand side through fast, efficient processes.

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- start supporting the demand side through fast, efficient processes.
- provide guaranteed supplementary funding, redeemable only for access to bio-databases.

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- data stamps

# Federal Funding of Bio-Databases

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- stop supporting the supply side of biodatabases through slow, inefficient processes.
- start supporting the demand side through fast, efficient processes.
- provide guaranteed supplementary funding, redeemable only for access to bio-databases.
- data stamps, AKA *food (for-thought) stamps* ?!

# Food (for thought) Stamps

---

## Funding Agencies could:

- provide a 10% supplement to **every** research grant in the form of “stamps” redeemable only at database providers.
- allow the “stamps” to be transferable among scientists, so that a market for them could emerge.
- provide funding only after the stamps have been redeemed at a database provider.

# Food (for thought) Stamps

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## Problems:

- how to estimate the amount of FFT stamps that would actually be redeemed (and thus the required budget set-aside).
- how to identify “approved” database providers.
- how to initiate the FFT system.
- etc etc

# Food (for thought) Stamps

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## **Alternatives (if no solution emerges):**

- increasingly inefficient research activities (abject failure will occur when it becomes simpler to repeat research than to obtain prior results).
- loss of access to bio-databases for public-sector research.
- movement of majority of “important” biological research into the private sector.
- loss of American pre-eminence (if other countries solve the problems first).